

## COMMUNICATIONS/ PUBLIC RELATIONS DEPARTMENT

Reference: AR Template

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# DEPARTMENT OF PAEDIATRICS AND CHILD HEALTH

Annual Report for Tygerberg Hospital and Stellenbosch University 2018

HEAD OF DEPARTMENT REPORT COMPILED BY

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#### **EXECUTIVE SUMMARY - 2018**

The year 2018 had posed many challenges as patient numbers were steadily increasing (respectively 6.5% for highly specialized paediatrics and 10.5% for general paediatrics), without the subsequent increase in staffing. Bed occupancy rate was especially high for general paediatrics (90%), neonatal ICU (86%) and paediatric ICU (86%). A further limitation was the lack of dedicated financial resources, which necessitated additional cost saving measures to be taken to ensure expenditure within allocated budget. Inborn survival rate of neonates was excellent with 98.5% of infants 500 grams and more surviving. Critical staff shortages include the lack of a second paediatric oncologist and a dedicated paediatric subspecialist for primary immunodeficiencies. Furthermore, without the assistance of the supernumerary recruited registrars (n=15) and senior registrars (n=5), we would not have been able to provide adequate service delivery - this is another major cost saving for Tygerberg hospital. The most important death causes remained septicaemia (16.9%) and pneumonia (10.5%), while the majority of patients that died were in the under-weight-forage category (12.1%). Major infrastructure improvements were funded by the Tygerberg included Children's hospital Trust and equipment the value R 672 866,35 and upgrading of the A9 intensive care complex to the value of R 193 400,70. The department successfully hosted the biannual South African Paediatric Association conference in August 2018 (chairs were Proff Ronald van Toorn and Pierre Goussard). Education programmes were successful with a high undergraduate pass rate (98%), and excellent postgraduate pass rate (12 completed Master's degrees) and an additional 6 postgraduate students were awarded their PhDs. The two research centres were successful in grant applications and contributed to the excellent number of research publications (n=132).

Professor Mariana Kruger

Part 1

## **RESOURCES AND OUTPUT**

## **Human Resources**

| Posts (full-time)                   | <u>Number</u>         | <u>Filled</u>           |
|-------------------------------------|-----------------------|-------------------------|
| Executive Head of Department/ Chief | 1                     | 1                       |
| Specialist                          |                       |                         |
| Head of Clinical Department/ Chief  | 1                     | 1                       |
| Specialist                          |                       |                         |
| Principal Specialists               | 5                     | 5                       |
| Senior Specialists                  | 25+1(5/8)             | 25+1(5/8)               |
| Senior Registrars                   | 9 (5 Supernumeraries) | 9 (5 Supernumeraries)   |
| *Registrars                         | 41 (including 15      | 41 (15 Supernumeraries) |
|                                     | Supernumeraries)      |                         |
| Medical Officers                    | 24                    | 24                      |
| Posts (sessional – hours per week)  |                       |                         |
| Specialist                          | 4 (52 hours)          | 4 (52 hours)            |
| Number of beds                      | 271                   | 271                     |

| Subspecialist Paediatrics | 2018    | 2017    | 2016     |
|---------------------------|---------|---------|----------|
| Inpatients                | 6269    | 5885    | 5658     |
|                           | (6.5%)  | (4.0%)  | (-4.1%)  |
| Outpatients               | 13694   | 13721   | 13224    |
|                           | (-0.1%) | (3.7%)  | (-1%)    |
| Summary of Output         |         |         |          |
| General Paediatrics       |         |         |          |
| Inpatients                | 10847   | 9810    | 9906     |
|                           | (10.5%) | (-0.9%) | (-11.8%) |
| Outpatients               | 12921   | 12637   | 11983    |
|                           | (2.2%)  | (5.4%)  | (-18%)   |

<sup>\*6</sup> registrars are appointed respectively Worcester (2), Paarl (2), KBH (1), KH (1)

## Output

## SUBSPECIALIST PAEDIATRICS

## Total patient admissions 2018 (Clinicom data)

| *A9<br>NICU | A9<br>PICU | A9<br>High Care | Trachea<br>Unit | <b>G1</b> | G3   | G9   | G10 | Total |
|-------------|------------|-----------------|-----------------|-----------|------|------|-----|-------|
| 509         | 803        | 209             | 51              | 1052      | 1064 | 1645 | 832 | 6165  |

<sup>\*</sup>A9 NICU includes High-care beds, since Clinicom cannot separate data.

## **Bed Occupancy Rate 2018(Clinicom data)**

| *A9  | A9   | A9        | Trachea | G1 | G3 | G9 | G10 | Total |
|------|------|-----------|---------|----|----|----|-----|-------|
| NICU | PICU | High Care | Unit    |    |    |    |     |       |

| 86% | 86% | 41% | 56% | 89% | 76% | 72% | 71% | 72% |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|
|-----|-----|-----|-----|-----|-----|-----|-----|-----|

<sup>\*</sup>A9 NICU includes High-care beds, since Clinicom cannot separate data. #BOR low due to nursing staff shortages.

## Neonatology

Staff: Prof J Smith, A Bekker, Drs S Holgate, G Kali, JCF du Preez, H Hassan, L Van Wyk, L Lloyd; 3 Senior Registrars (2 supernumerary) – Dr N Paulse, Dr MW Kariuki, Dr K Onwona-Agyeman; 6 Registrars; 7 Medical Officers

## **A9 Neonatal Intensive Care Unit**

Staff: 1 Consultant (on rotation), 2 Registrars, 1 Medical Officer

| Beds n=8                  | 2018     | 2017        | 2016       |
|---------------------------|----------|-------------|------------|
| Admissions                | 510      | 461         | 442        |
| Average NICU Stay in Days | 8        | 8           | 9          |
| Average Bed Occupancy %   | 86%      | 81%         | 88%        |
| % Growth                  | +10.6%   | +4.3%       | -12.6%     |
| Deaths                    | 97 (19%) | 108 (23.4%) | 76 (17.1%) |

- Referrals from outside TCH (outborns) (n) = 178; 34.9% of admissions
- Top 5 referring hospitals:
  - o KBH 67
  - o Paarl 62
  - o HH 37
  - o Worcester 22
  - o KH 4
- Referrals for therapeutic hypothermia (TH): total 108 (21.2% of admissions)
  - o Inborn babies treated with therapeutic hypothermia (n) = 47
  - Referrals from referral hospitals (n) = 61: (75.4% of outborn referrals from HH & KBH)
    - 16 from HH
    - 30 from KBH
    - 13 from Paarl
- Surgical admissions: 92 (18.04% of admissions)
- Cranial ultrasounds: ±950
- Neonatal echocardiography: ±150

## Ward G1: Neonatal Unit - Babies born outside TBH

Staff: 2 Consultants (on rotation), 2 Registrar, 1 Medical Officers, 1 Intern

| Beds n=30                 | 2018    | 2017      | 2016  |
|---------------------------|---------|-----------|-------|
| Admissions                | 1052    | 989       | 1115  |
| Average Ward Stay in Days | 9       | 10        | 9     |
| Average Bed Occupancy %   | 89%     | 85%       | 91%   |
| % Growth                  | +6.4%   | -11.3%    | -3.2% |
| Deaths                    | 21 (2%) | 22 (2.2%) | 36    |

## **Highly Specialized Paediatrics**

## **A9 Paediatric Intensive Care Unit**

Staff: Prof P Goussard, Drs N Parker, Dr I Appel, 3 Registrars, Medical Officer

| Beds <i>n=10</i>              | 2018      | 2017       | 2016  |
|-------------------------------|-----------|------------|-------|
| Admissions                    | 818       | 728        | 745   |
| Average Hospital Stay in Days | 4         | 4          | 4     |
| Average Bed Occupancy %       | 86%       | 84%        | 81.6% |
| % Growth                      | +10.3%    | +10.1%     | +4.6% |
| Deaths                        | 65 (7.9%) | 79 (10.9%) | 62    |

## Paediatric Intensive Care Unit (PICU) bookings and admitted for elective postoperative beds 2018 (Table 1 and 2)

- Total bookings: n=372
  - o Elective admissions: n=225 (60.5% of bookings and 27.5% of total PICU admissions)
    - Paediatric surgery: 26.7%
    - Thoracic surgery: 19.6%
    - Ear Nose and Throat (ENT): 15.6%
    - Other (as detailed in Table 1)
  - Patients with booking but not admitted to PICU: 39.5% (n=147)
    - 33.3% did not need a bed after surgery (n=49)
    - Cancellations of surgery: 22.4% (n=33)
    - Unknown reasons: 17% (n=25)
    - PICU refusal as patients not deemed to need intensive care: n=2
       (Ophthalmology and MRI)
    - Bed not available: 15.6% (n=23) especially during the months March to July 2018.
      - Paediatric surgery: n=7
      - ENT: n=5
      - Other reasons: nursing shortage n=2; CRE n=3; MRSA n=1

Table 1. Outcome of all Bookings for elective post-op beds in PICU (number / %)

| Surgical<br>discipline | Admit-<br>ted | NA* - not<br>required<br>adm to<br>surgical /<br>medical<br>ward<br>postop | NA* -<br>bed<br>cancelled<br>by<br>surgeon | Surgery<br>cancelled | NA –<br>PICU<br>Bed<br>NA | NA –<br>PICU<br>bed<br>cancelle<br>d by<br>anaesth<br>etist | Bed refused by PICU - not required (adm to ward post op) | Not<br>contacte<br>d for<br>bed<br>space | Nursing<br>short-<br>staffed | PICU<br>bed<br>NA* -<br>CRE** | PICU<br>bed<br>NA –<br>MRSA*<br>** |
|------------------------|---------------|----------------------------------------------------------------------------|--------------------------------------------|----------------------|---------------------------|-------------------------------------------------------------|----------------------------------------------------------|------------------------------------------|------------------------------|-------------------------------|------------------------------------|
| Thoracic               | 44<br>(19.6%) | 6                                                                          | 1                                          | 6                    | 1                         |                                                             |                                                          | 2                                        | 1                            |                               |                                    |
| General                | 60<br>(26.7%) | 13                                                                         | 4                                          | 12                   | 4                         |                                                             |                                                          | 7                                        |                              | 3                             |                                    |
| ENT                    | 35<br>(15.6%) | 5                                                                          | 4                                          | 10                   | 4                         |                                                             |                                                          | 8                                        |                              |                               | 1                                  |
| Orthopaedic            | 7 (3.1%)      | 1                                                                          |                                            |                      | 1                         | 1                                                           |                                                          |                                          |                              |                               |                                    |
| Spinal                 | 25<br>(11.1%) |                                                                            |                                            | 3                    | 2                         |                                                             |                                                          |                                          |                              |                               |                                    |
| Plastics               | 10<br>(4.4%)  | 2                                                                          |                                            | 1                    | 2                         |                                                             |                                                          | 2                                        |                              |                               |                                    |

| Urology                        | 19                 | 6          | 2            |              | 2                | 1        |          | 2            |         |                 |                 |
|--------------------------------|--------------------|------------|--------------|--------------|------------------|----------|----------|--------------|---------|-----------------|-----------------|
| Post CT /<br>MRI bed<br>booked | (8.4%)<br>3 (1.3%) | 4          |              |              |                  |          | 1        | 1            |         |                 |                 |
| Ophthalmol ogy                 | 3 (1.3%)           |            | 1            |              |                  |          | 1        | 1            |         |                 |                 |
| Neurosurger<br>y               | 16<br>(7.1%)       | 7          | 1            | 1            | 1                |          |          | 2            | 1       |                 |                 |
| Post<br>Bronchos-<br>copy      | 3 (1.3%)           | 5          |              |              |                  |          |          |              |         |                 |                 |
| Total 372                      | 225<br>(60.5%)     | 49 (13.2%) | 13<br>(3.5%) | 33<br>(8.9%) | 17<br>(4.6<br>%) | 2 (0.5%) | 2 (0.5%) | 25<br>(6.7%) | 2(0.5%) | 3<br>(0.8<br>%) | 1<br>(0.3<br>%) |

<sup>\*</sup>NA Patients booked for post-op bed and not admitted to PICU

Table 2. Patients booked postoperatively for PICU bed and not admitted to PICU (number/%)

| Surgical<br>discipline         | Admitted<br>to surgical<br>/ medical<br>ward<br>postop | PICU bed<br>cancelled<br>by surgeon | Not<br>contacted<br>for PICU<br>bed | Surgery<br>cancelled | PICU Bed<br>not<br>available | PICU bed<br>cancelled<br>by<br>anaestheti<br>st | Bed<br>refused by<br>PICU - not<br>required |
|--------------------------------|--------------------------------------------------------|-------------------------------------|-------------------------------------|----------------------|------------------------------|-------------------------------------------------|---------------------------------------------|
| Thoracic                       | 6                                                      | 1                                   | 2                                   | 6                    | 2                            |                                                 |                                             |
| General                        | 13                                                     | 4                                   | 7                                   | 12                   | 7                            |                                                 |                                             |
| ENT                            | 5                                                      | 4                                   | 8                                   | 10                   | 5                            |                                                 |                                             |
| Orthopaedic                    | 1                                                      |                                     |                                     |                      | 1                            | 1                                               |                                             |
| Spinal                         |                                                        |                                     |                                     | 3                    | 2                            |                                                 |                                             |
| Plastics                       | 2                                                      |                                     | 2                                   | 1                    | 2                            |                                                 |                                             |
| Urology                        | 6                                                      | 2                                   | 2                                   |                      | 2                            | 1                                               |                                             |
| Post CT /<br>MRI bed<br>booked | 4                                                      |                                     | 1                                   |                      |                              |                                                 | 1                                           |
| Ophthalmo-<br>logy             |                                                        | 1                                   | 1                                   |                      |                              |                                                 | 1                                           |
| Neurosurger<br>y               | 7                                                      | 1                                   | 2                                   | 1                    | 2                            |                                                 |                                             |
| Post<br>Bronchos-<br>copy      | 5                                                      |                                     |                                     |                      |                              |                                                 |                                             |
| Total 147                      | 49 (33.3%                                              | 13 (8.8%)                           | 25 (17.0%)                          | 33 (22.4%)           | 23 (15.6%)                   | 2 (1.4%)                                        | 2 (1.4%)                                    |

<sup>\*\*</sup>CRE Carbapenem resistant Enterobacteriaceae

<sup>\*\*\*</sup>MRSA Methicillin resistant Staphylococcus Aureus

## A9 Paediatric High Care Unit

Staff as mentioned above.

| Beds n=4                      | 2018  | 2017  | 2016 |
|-------------------------------|-------|-------|------|
| Admissions                    | 209   | 181   | 108  |
| Average Hospital Stay in Days | 3     | 4     | 3.2  |
| Average Bed Occupancy %       | 41%   | 34%*  | 25%  |
| Growth %                      | 15.4% | 67.5% |      |
| Deaths                        | 2     | 2     | 3    |

<sup>\*</sup> Due to nursing staff shortages

## **A9 Tracheostomy Unit**

Staff as mentioned above:

| Beds n=6                | 2018 | 2017  | 2016 |
|-------------------------|------|-------|------|
| Admissions              | 51   | 109   | 96   |
| Average Bed Occupancy % | 56%  | 62%   | 44%  |
| Growth %                | -53% | 13.5% |      |
| Deaths                  | 1    | 1     | 0    |
| Decannulation           |      | 10    | /    |

## Ward G9 Paediatric Pulmonology and Allergy\*

Staff: Prof P Goussard, Dr J Morrison, 2 Senior Registrars (Dr L Green & Dr Y Mulambia), 2 Registrars, Shared Medical Officer in G9. \*Allergy (largely outpatients service) \*Prof S Kling, \*Prof E Zöllner, \*Dr Y Kooblal (4hrs per week)

| Pulmonology Beds n=10         | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions Pulmonology        | 995  | 956  | 977  |
| Average Hospital Stay in Days | 5.8  | 4.5  | 5.6  |
| Average Bed Occupancy %       | 65%  | 93%  | 94%  |
| % Growth                      | +4%  | -2%  | 9%   |
| Deaths                        | 1    | 1    | 2    |

<sup>\*</sup>Doctors attached to other services, but responsible for allergy clinics

## Theatre procedures and Other Activities

Bronchoscopies: 465Thoracic surgery: 63

• 6 Outreach clinical teaching sessions and ward rounds to Karl Bremer hospital

## **Ward G9 Neurology**

Staff: Prof R van Toorn, Prof R Solomons, Dr P Springer, 1 Senior Registrar (Dr T Katangwe), Dr A Thomas(8hrs per week), Dr Y Kooblal(4hrs per week) 2 Registrars

| Beds <i>n=10</i>              | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions                    | 501  | 421  | 388  |
| Average Hospital Stay in Days | 6    | 6.0  | 6.1  |
| Referral other wards          | 740  | 680  | 550  |

| Average Bed Occupancy % | 76%  | 72%    | 65%   |
|-------------------------|------|--------|-------|
| % Growth                | +19% | +10.8% | -8.8% |
| Deaths                  | 8    | 4      | 5     |

## **Other Activities**

Paediatric neurology consultations in other wards/PICU/NICU: 245

• Paediatric EEG from Worcester hospital: 134

• Paediatric and neonatal EEG's from TBH reported by the 2 consultants: 480

6 Outreach clinics to Paarl hospital (patients seen): 95
4 Outreach clinics Worcester hospital (patients seen): 105
2 Clinics at Alta du Toit special school (patients seen): 40

## **Ward G9 Paediatric Endocrinology**

Staff: Prof E Zöllner, Dr M Grantham (25 hrs per week), Registrar, and shared intern for G9

| Beds n=5                                | 2018              | 2017              | 2016                        |
|-----------------------------------------|-------------------|-------------------|-----------------------------|
|                                         | 102               | 120               | 95                          |
| Patients admitted                       | Diabetic          | Diabetic          |                             |
| Patients admitted                       | 150               | 131               | 167                         |
|                                         | Endocrinology     | Endocrinology     |                             |
| Admission Subtotal                      | 252               | 251               | 262                         |
| Admissions other wards, day             | 8                 | 10                | 14                          |
| cases                                   |                   |                   |                             |
| Admissions total                        | 260               | 261               | 275                         |
| Ward reviews                            | 14                | 6                 | 98                          |
| Average Hospital Stay in Days           | 6                 | 6                 | 6                           |
| Average Bed Occupancy %                 | 88% (not counting | 81% (not counting | 84% (not counting outliers) |
|                                         | outliers)         | outliers)         | (                           |
| % Growth                                | N/A               | -5%               | 17% (not counting ward      |
| ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,, |                   |                   | reviews)                    |
| Deaths                                  | 0                 | 0                 | 0                           |

Endocrine tests performed on; 53 patients

## **Ward G10 Gastroenterology**

Staff: Prof ED Nel, 2 Senior Registrars (Dr LT Radebe & Dr U Chowdhury), Registrar, and 1 Intern.

| Beds n=9                      | 2018 | 2017 | 2016     |
|-------------------------------|------|------|----------|
| Admissions                    | 262  | 254  | 185      |
| Average Hospital Stay in Days | 7    | 8    | 11       |
| Average Bed Occupancy %       | 70%  | 82%  | 75%      |
| % Growth                      | +3%  | +37% | -19%     |
| Deaths                        | 7    | 4    | 5 (2,7%) |

## Theatre procedures

Paediatric Endoscopy: 47

Adult Colonoscopy: 281

• Other endoscopic procedures: 445

## **Ward G10 Infectious Diseases Unit**

Staff: Prof MF Cotton, Prof H Rabie (HIV Clinic), Dr P Rose, 2 Registrars

| Beds <i>n=14</i>              | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions                    | 374  | 353  | 269  |
| Average Hospital Stay in Days | 9    | 9    | 14   |
| Average Bed Occupancy %*      | 77%  | 82%  | 75%  |
| % Growth                      | +6%  | +31% | -15% |
| Deaths                        | 14   | 4    | 2    |

<sup>\*</sup> Clinicom data combined for gastroenterology, cardiology and infectious diseases

• 10 new referrals per week.

• Telephonic referrals: 100 per week (including outside calls)

• Stewardship rounds: 2 per week.

Transfers out: 17Transfers in: 50

## **G10 Cardiology**

Cross Platform Staff:

Tygerberg Hospital (TBH): Prof J Lawrenson, Dr B Fourie, 2 Registrars

Red Cross War Memorial Children's Hospital (RXH): 2 Consultants, 2 Senior Registrar

| Beds n=10 (G3=5, G10=5)       | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions                    | 159  | 146  | 171  |
| Average Hospital Stay in Days | *    | *    | 7    |
| Average Bed Occupancy %       | *    | *    | *    |
| Inpatient Echocardiography    | 923  | 845  | 836  |
| Inpatient Consultations       | 945  | 882  | 875  |
| Growth numbers                | 63   | 7    |      |
| Growth %                      | 7.1% | 0.8% | /    |
| Deaths                        | 2    | 2    | 3    |

<sup>\*</sup> Clinicom data combined for gastroenterology, cardiology and infectious diseases

Worcester Clinic: 164 Paarl Clinic: 132

## Ward G3 Oncology & Haematology

Staff: Prof M Kruger, Drs A van Zyl, R Uys, 1 Senior Registrar (Dr Lillian Gesami-Steytler, later replaced by Dr L Schoonraad), 2 Registrars

| Beds n=9     | 2018 | 2017 | 2016 | 2015 |
|--------------|------|------|------|------|
| New patients |      |      |      |      |
| Haematology  | 44   | 46   | 43   | 42   |

| New patients<br>Oncology                          | 70             | 60             | 72      | 73    |
|---------------------------------------------------|----------------|----------------|---------|-------|
| New patients Total                                | 114            | 106            | 115     | 114   |
| Admissions                                        | 783            | 734            | 759     | 678   |
| % Growth                                          | +6.7%          | -3.2%          | 11.4%   | 11.9% |
| Average Hospital Stay in Days                     | 5.25           | 5.25           | 6.05    | 4.9   |
| Average Bed<br>Occupancy %                        | 123%           | 120.5%         | 148.25% | 93.8% |
| Total deaths in G3                                | 12             | 12             | 14      | 9     |
| Expected palliative                               | 8              | 8              |         |       |
| Sepsis/other                                      | 4              | *4             |         |       |
| Referrals                                         | Haematology 70 | Haematology 73 |         |       |
| G3 Day Patients<br>(Haem/Onco)=<br>Ward attenders | 1698           | 2094           | 2806    | 1905  |
| Chemotherapy administrations                      | 2051           | 2044           | 2590    | 1944  |
| Bone marrow aspirations and biopsies              | 101            | 108            | 143     | 122   |
| Intra-thecal chemotherapy procedures              | 309            | 396            | 263     | 286   |

<sup>\*</sup>G3Ward data

## Ward G3 Rheumatology & Immunology

Staff: Prof M Esser, Dr D Abraham (15 hrs per week), Registrar Shared with G3, Shared Medical Officer for G3

| Beds n=4                      | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions                    | 120  | 110  | 90   |
| (Ward attenders)              | 130  | (82) |      |
| Average Hospital Stay in days | 5    | 2    | 3    |
| Average bed occupancy %       | 36%  | 16%  |      |
| % Growth                      | 18%  | 22%  | 36%  |
| G3 Day Patients (Rheuma&Imm)  | 94   | 6    |      |
| Deaths                        | 0    | 0    | 0    |

## **Ward G3 Nephrology**

Staff: Dr CJ du Buisson, Dr JL Shires – part-time, Registrar shared with haematology & Immunology, Shared Intern for G3

| Beds n=5 | 2018 | 2017 | 2016 |
|----------|------|------|------|
|----------|------|------|------|

| Admissions                      | 138          | 120         | 134          |
|---------------------------------|--------------|-------------|--------------|
| Average Hospital Stay in days   | 5            | 5           | 5            |
| Average bed occupancy %         | 41%***       | 31% ***     | 45%          |
| % Growth                        | 15%          | -32% ***    | -44%         |
| Deaths                          | 1            | 0           |              |
| Renal Biopsies*                 | 22           | 23          | 21           |
| Acute Dialysis**                | 16           | 11          | 12           |
| Panal nationts outside of C2*** | 276          | 384         | 354          |
| Renal patients outside of G3*** | 28% decrease | 9% increase | 26% increase |
| Referrals****                   | 431          | 428         | 345          |

- \* Done in C4B Theatre under general anaesthesia
- \*\* All done in PICU and NICU, 1 patient was dialysed in G3 for 5weeks
- \*\*\* Double the amount of nephrology patients are admitted outside of G3. 42% are in Neonatal wards (postnatal or antenatal diagnosis), 33% in G7 and 25% in G10 (due to infectious infections or lack of space in G3). 40% of admissions to G7 and G10 is due to a lack of space in G3.
- \*\*\*\* These are patients on which only advice needed in the acute phase, not continuous care and referred to outpatient service for further treatment. 91 of these pateints where seen on a regular basis in G4 as part of our cross disciplinary partnership with urology.
- \*\*\*\* These are patients on which only advise needed in the acute phase, not continuous care and referred to outpatient service for further treatment

#### **OUTPATIENT COMPLEX**

## **Subspecialist Clinics**

|                                       | 2018  | 2017  | 2016  |
|---------------------------------------|-------|-------|-------|
| Clinics                               | Total | Total | Total |
| Haematology                           | 306   | 285   | 307   |
| Immunology                            | 169   | 229   | 191   |
| Oncology                              | 484   | 471   | 361   |
| Pulmonology                           | 1194  | 1130  | 1063  |
| Gastroenterology                      | 635   | 581   | 563   |
| High-risk Babies                      | 2018  | 2056  | 2005  |
| Neurology & Developmental paediatrics | 2375  | 2176  | 2185  |
| Allergy                               | 566   | 550   | 580   |
| Premature Follow-up                   | 210   | 262   | 234   |
| Nephrology*                           | 1547  | 1482  | 1469  |
| Cardiology                            | 1099  | 1108  | 1073  |
| Diabetic                              | 742   | 720   | 703   |
| Endocrinology                         | 685   | 666   | 689   |
| Rheumatology                          | 359   | 367   | 348   |
| Infectious diseases                   |       |       |       |
| ТВ                                    | **752 | 855   | 890   |
| HIV                                   | 473   | 467   |       |
| Genetics                              | 468   | 529   | 393   |
| Pharmacy prescriptions                | 153   | 254   | 170   |
| Total                                 | 13694 | 13721 | 13224 |

<sup>\*</sup>Centre for Referral for Congenital Anomalies of the Kidney and Urinary Tract.

## \*\*68 Patients seen in GGround due to limited number to be seen in C3A Neuro-development:

- Developmental Medicine functions largely as an outpatient service:
  - o 8 Clinics at Paarl School (patients seen): 80
  - o Autism Diagnostic Observations Schedule (ADOS) evaluation: 14 children
  - o Griffith Developmental Scales assessments: 20.

#### HIV service:

• Total new patients initiating ART: 77 children (46 infants <12 months)

## **GENERAL SPECIALIST SERVICES**

## **Total Patient Admissions 2018 (Clinicom data)**

| GG<br>Short Stay | G2   | G7   | G8   | J3  | Total  |
|------------------|------|------|------|-----|--------|
| 5306             | 2072 | 1308 | 1169 | 992 | 10 847 |

## **Bed Occupancy Rate 2018 (Clinicom data)**

| GG<br>Short Stay | G2  | G7  | G8  | J3   | Total |
|------------------|-----|-----|-----|------|-------|
| 79%              | 94% | 78% | 90% | 108% | 90%   |

## **Neonatology**

## Ward G2 Neonatal Unit - Babies born in TBH

Staff: 2 Consultants (on rotation), 1 Registrar, 3 Medical Officers, 1 Intern (if available)

| Beds <i>n=27</i>          | 2018      | 2017      | 2016      |
|---------------------------|-----------|-----------|-----------|
| Admissions                | 2072      | 2142      | 2087      |
| Average Ward Stay in Days | 4         | 5         | 5         |
| Average Bed Occupancy     | 94%       | 105%      | 104%      |
| % Growth                  | -3.2%     | +2.6%     | 0.96%     |
| Deaths                    | 71 (3.4%) | 70 (3.3%) | 94 (4.5%) |

## Ward J3 Neonatal Unit – Babies born in TBH

Staff: 1 Consultant (on rotation), 1 Registrar, 1 Medical Officer, 1 Intern

| Beds <i>n=25</i>          | 2018  | 2017  | 2016  |
|---------------------------|-------|-------|-------|
| Admissions                | 992   | 908   | 972   |
| Average Ward Stay in Days | 10    | 10    | 10    |
| Average Bed Occupancy     | 108%  | 101%  | 109%  |
| % Growth                  | +9.3% | -6.6% | -3.4% |

| Deaths | 3 (0.3%)  | 2 (0.2%)  | 1(0.1%)  |
|--------|-----------|-----------|----------|
| Deaths | 3 (0.370) | 2 (0.2/0) | 1(0.1/0) |

## Ward G8 Low Care Neonates & Kangaroo-mother Care - Step-down Facility

Staff: 1 Consultant (on rotation), 1 Medical Officer, 2 Interns

| Beds <i>n=30</i>          | 2018     | 2017   | 2016     |
|---------------------------|----------|--------|----------|
| Admissions                | 1169     | 1057   | 1013     |
| Average Ward Stay in Days | 8        | 9      | 10       |
| Average bed occupancy %   | 90%      | 87%    | 88%      |
| % Growth                  | +9.2%    | +4.3%  | 10.4%    |
| Deaths                    | 4 (3.2%) | 0 (0%) | 3 (0.3%) |

## Postnatal wards (J5, J2, C2A)

Staff: 1 intern from neonatal pool

3640 clinical contacts/year

## **Short-stay G Ground: <48-hour Admissions**

Staff: Prof S Schaaf, Drs E Malek, L Smit, A Redfern, 2 Registrars, (1 Emergency Medicine Registrar), 2 Medical Officers, 2 Interns

| Beds n=24                     | 2018 | 2017 | 2016 |
|-------------------------------|------|------|------|
| Admissions                    | 5306 | 4567 | 4643 |
| Average Hospital Stay in Days | 1    | 1    | 1    |
| Average Bed Occupancy %       | 79%  | 63%  | 62%  |
| % Growth                      | +16% | -2%  | -22% |
| Average Admissions per day    | 15   | 13   | 13   |
| Deaths                        | 11   | 7    | 16   |

## **Ward G7 General Paediatrics**

Staff: Prof S Kling, Dr H Finlayson, 1 Registrar, 2 Medical Officers, 2 Interns

| Beds <i>n=25</i>              | 2018 | 2017      | 2016  |
|-------------------------------|------|-----------|-------|
| Admissions                    | 1308 | 1136      | 1191  |
| Admissions to HCU             | 441  | 134 (6mo) |       |
| Average Hospital Stay in Days | 6    | 6         | 6     |
| Average Bed occupancy         | 78%  | 74%       | 77%   |
| Increase in Admissions        | +15% | -4.3%     | -3.3% |
| Deaths                        | 15   | 10        | 17    |

<sup>\*</sup>Clinicom data

**General Paediatrics: Emergency & Clinics** 

| Clinics | 2018 | 2017 | 2016 |
|---------|------|------|------|
|         |      |      |      |

| OPD 8am-4pm:<br>Emergency & Routine                     | 8 282 (GG total) | 8 017 (GG total)<br>1203 OPD<br>6814 Emergency | 7 661 (GG total)<br>1643 OPD<br>6018 Emergency |
|---------------------------------------------------------|------------------|------------------------------------------------|------------------------------------------------|
| Daily average seen                                      | 33               | 32                                             | 31                                             |
| Annual after hours:<br>4pm-8am & weekend<br>– Emergency | 4 639            | 4 620                                          | 4 322                                          |
| Daily average seen<br>after hours (4pm-<br>8am)         | 13               | 13                                             | 12                                             |
| Emergencies after hours: 4pm-8am %                      | 36%              | 36%                                            | 36%                                            |
| Total 12 921                                            |                  | 12 637                                         | 11 983                                         |
| Increase                                                | +2%              | +5%                                            |                                                |

## 2018 Morbidity & Mortality

#### PPIP data from PPIP database 2018

Total inborn births in TCH (TBH): = 8068

(Stillbirths = 541) (Total live births = 7527)

## **Inborn mortality rates at TCH:**

• Perinatal Mortality Rate (PNMR) (≥500g) = 77.2/1000 • Perinatal Mortality Rate (PNMR) (≥1000g) = 44.1/1000 • Early Neonatal Death Rate (ENNDR) (≥500g) = 10.9/1000 • Early Neonatal Death Rate (ENNDR) (≥1000g) = 5.1/1000 • Late Neonatal Death Rate (LNNDR) (≥500g) = 4.1/1000 • Late Neonatal Death Rate (LNNDR) (≥1000g) = 1.9/1000

## **Inborn Survival Rate for babies born alive at TCH:**

Survival Rate (≥500g) = 98.5%
 Survival Rate (≥1000g) = 99.3%

## Inborn mortality of babies born alive per birth weight category:

| Birth Weight  | % Mortality 2018 | % Mortality 2017      | % Mortality 2016 |  |
|---------------|------------------|-----------------------|------------------|--|
| < 1000g       | 22.1             | <b>26.3</b> (58/220)  | 27.6             |  |
| 1000 to 1499g | 3.6              | <b>6.5</b> (31/471)   | 5.6              |  |
| 1500 to 1999g | 1.6              | <b>2.5</b> (18/722)   | 2.3              |  |
| 2000 to 2499g | 0.4              | <b>1.1</b> (12/1124)  | 0.9              |  |
| ≥2500g        | 0.3              | <b>0.56</b> (28/4928) | 0.2              |  |

Mortality % calculated for each weight category as: number of deaths per number of live births in that weight category for 2018.

## Causes of death of babies born in TCH (TBH):

Total inborn neonatal deaths on PPIP = 113

| Cause of Death       | % of Total 2018* | % of Total 2017 | % of Total 2016 |  |
|----------------------|------------------|-----------------|-----------------|--|
| Prematurity-related  | 49.4             | 44.4            | 61.4            |  |
| Complications        |                  |                 |                 |  |
| Infection-related    | 15.3             | 12.9            | 10.5            |  |
| Peripartum Hypoxia   | 15.3             | 16.9            | 7.9             |  |
| Congenital Anomalies | 12.9             | 19.4            | 14.9            |  |
| Other                | 7.1              | 6.4             | 5.3             |  |

<sup>\*</sup>Reflects estimated figures as only ~80% of data complete, and is thus not fully validated yet.

## % Deaths of babies referred to TCH per weight category:

Total neonatal deaths in babies referred to TCH = 45

| Birth Weight    | % Mortality 2018<br>(n= 45) | % Mortality 2017<br>(n=62) | % Mortality 2016<br>(n= 56) |
|-----------------|-----------------------------|----------------------------|-----------------------------|
| <1000g          | 11.1% (5/45)                | 7.8 (4/51)                 | 25                          |
| 1000 to 1499g   | 22.2% (10/45)               | 17.6 (9/51)                | 35                          |
| 1500 to 1999g   | 2.2% (1/45)                 | 17.6 (9/51)                | 10                          |
| 2000 to 2499    | 13.3% (6/45)                | 7.8 (4/51)                 | 7.5                         |
| ≥2500g          | 28.9% (13/45)               | 49 (25/51)                 | 25                          |
| Wt not recorded | 22.2% (10/45)               |                            |                             |

<sup>%</sup> calculated as the number of referred (outborn) deaths in each weight category out of all the referred (outborn) neonatal deaths (all weight categories).

## Outborn babies mortality (total numbers) according to referral area of origin:

| Geographic Service Area            | 2018 (n= 45) | 2017 (n=62) | 2016 (n=56) |
|------------------------------------|--------------|-------------|-------------|
| Metro East                         | 17           | 30          | 26          |
| Winelands (Paarl area)             | 15           | 11          | 14          |
| Overberg (Worcester drainage area) | 6            | 10          | 5           |
| Helderberg                         | 5            |             |             |
| Khayelitsha                        | 2            |             |             |
| Unknown                            |              | 0           | 17          |

## CHPIP MORTALITY for all sub-specialist paediatric wards (2018)

**Table 1**: All deaths per ward and age as on Clinicom excluding children>29 days of age dying in the neonatal deaths

| Ward     | Total       | 0-7<br>days | 8-28<br>days | 29 days -<br>11months | 12 -59<br>months | >5-years |
|----------|-------------|-------------|--------------|-----------------------|------------------|----------|
| PICU     | 66 (48.89%) | 1           | 4            | 31                    | 17               | 13       |
| PICU -HC | 4 (0.03%)   |             |              | 3                     |                  |          |

| G10   | 14 (10.37%) | 0            | 0            | 6              | 5              | 3              |
|-------|-------------|--------------|--------------|----------------|----------------|----------------|
| G9    | 11 (8.15%)  | 0            | 0            | 4              | 3              | 4              |
| G3    | 13 (9.63%)  | 0            | 0            | 1              | 5              | 7              |
| G7    | 16 (11.85%) | 0            | 2            | 5              | 8              | 1              |
| GG    | 11 (8.15%)  | 0            | 0            | 3              | 5              | 3              |
| Total | 135*        | 1<br>(0.74%) | 6<br>(4.44%) | 53<br>(39.26%) | 43<br>(31.85%) | 32<br>(23.70%) |

Median duration of stay at death 4 days IQR 1 – 14 ) [mean 11 days]

Table 2: Audited deaths per age and firm as per Child PIP

| Ward             | 0 - 7<br>days | 8 - 28<br>days | 28 days -<br>1 Year | 1 - 5<br>years | 5 - 13<br>years | 13 – 18<br>Years | Unknown | Total   |
|------------------|---------------|----------------|---------------------|----------------|-----------------|------------------|---------|---------|
| A9 PICU          | 0             | 4              | 28                  | 17             | 15              | 2                | 0       | 66      |
| Oncology         | 0             | 0              | 0                   | 5              | 6               | 0                | 1       | 12      |
| Gastroenterology | 0             | 0              | 5                   | 3              | 0               | 0                | 0       | 8       |
| Neurology        | 1             | 0              | 0                   | 2              | 2               | 0                | 0       | 5       |
| ID               | 0             | 0              | 1                   | 0              | 0               | 1                | 2       | 4       |
| Neonatology*     | 0             | 0              | 3                   | 0              | 0               | 0                | 0       | 3       |
| Cardiology       | 0             | 0              | 0                   | 2              | 0               | 0                | 0       | 2       |
| Nephrology       | 0             | 0              | 1                   | 0              | 0               | 0                | 0       | 1       |
| G7               | 0             | 1              | 5                   | 5              | 1               | 0                | 0       | 12      |
| GGround          | 0             | 1              | 4                   | 3              | 3               | 0                | 0       | 11      |
| Total            | 1             | 6              | 47                  | 37             | 27              | 3                | 3       | 124     |
|                  |               |                |                     |                |                 |                  |         | 58      |
| Girls            | 1             | 1              | 20                  | 17             | 14              | 2                | 3       | (47,8%) |
|                  |               |                |                     |                |                 |                  |         | 66      |
| Boys             | 0             | 5              | 27                  | 20             | 13              | 1                | 0       | (53,2%) |

<sup>\*22</sup> infants older than 28 days of age died in neonatal service (only 3 audited deaths)

Table 3: Length of stay as per Child PIP

| Length of stay | Number | Percent |
|----------------|--------|---------|
| DOA            | 4      | 3,2     |
| < 24 Hours     | 33     | 26,6    |
| 1-3 Days       | 27     | 21,8    |
| 4-7 Days       | 21     | 16,9    |
| 8-14 Days      | 19     | 15,3    |
| > 14 Days      | 20     | 16,1    |
| Unknown        | 0      | 0       |

Table 4:

| Main Cause Of Death | Number | % |
|---------------------|--------|---|
|---------------------|--------|---|

<sup>\*</sup>Excludes children that died in neonatal service

| Septicaemia, possible serious bacterial infection           | 21 | 16,9 |
|-------------------------------------------------------------|----|------|
| Pneumonia, ARI                                              | 13 | 10,5 |
| Acute diarrhoea, hypovolaemic shock                         | 10 | 8,1  |
| Other Respiratory System (specify)                          | 8  | 6,5  |
| Other Nervous System (specify)                              | 8  | 6,5  |
| Heart failure, Pulmonary Oedema                             | 7  | 5,6  |
| Other diagnosis (specify)                                   | 6  | 4,8  |
| Tumours                                                     | 6  | 4,8  |
| Status epilepticus                                          | 5  | 4    |
| Other Digestive System (specify)                            | 5  | 4    |
| Other Circulatory System (specify)                          | 4  | 3,2  |
| TB: Meningitis                                              | 3  | 2,4  |
| Cirrhosis, Portal Hypertension, Liver Failure,<br>Hepatitis | 3  | 2,4  |
| Myocarditis                                                 | 2  | 1,6  |
| Other accidents (incl. Drowning; specify)                   | 2  | 1,6  |
| Acute renal failure                                         | 2  | 1,6  |
| Cardiomyopathy                                              | 2  | 1,6  |
| Hospital-acquired infection                                 | 2  | 1,6  |
| Other possible serious infection (specify)                  | 2  | 1,6  |
| Other Endocr, Nutritional, Metabol. (specify)               | 1  | 0,8  |
| Other Oncology/Haematology (specify)                        | 1  | 0,8  |
| Other Poisoning (specify)                                   | 1  | 0,8  |
| TB: Miliary, other extra-pulmonary                          | 1  | 0,8  |
| Surgical (Appendix, hernia, intestines, peritoneum)         | 1  | 0,8  |
| Hypoglycaemia                                               | 1  | 0,8  |
| IDDM, DKA                                                   | 1  | 0,8  |
| III-defined/Unknown cause of mortality                      | 1  | 0,8  |
| Leukaemias                                                  | 1  | 0,8  |
| Meningitis: bacterial                                       | 1  | 0,8  |
| Meningitis: viral (meningo-encephalitis)                    | 1  | 0,8  |
| Chronic renal disease                                       | 1  | 0,8  |
| Congenital Heart Disease                                    | 1  | 0,8  |

## Table 5:

| Weight Category | Number | %    |
|-----------------|--------|------|
| 1. OWFA         | 3      | 2,4  |
| 2. Normal       | 71     | 57,3 |
| 3. UWFA         | 15     | 12,1 |
| 4. Marasmus     | 8      | 6,5  |
| 5. Kwashiorkor  | 10     | 8,1  |

| 6. M-K (Marasmic-Kwashiorkor) | 11 | 8,9 |
|-------------------------------|----|-----|
| Unknown                       | 6  | 4,8 |

 Table 6: Death by HIV status

| HIV status | Number |
|------------|--------|
| Negative   | 77     |
| Unknown    | 16     |
| Exposed    | 13     |
| Not tested | 12     |
| Infected   | 4      |
| No result  | 2      |

 Table 7: Referring facilities

| District                                                      | Subdistrict                               | Number | %    |
|---------------------------------------------------------------|-------------------------------------------|--------|------|
| Unknown                                                       | Unknown                                   | 58     | 46,8 |
| Cape Winelands District                                       | Breede River/Winelands Local Municipality | 7      | 5,6  |
| Municipality (13=10,5%)                                       | Drakenstein Local Municipality            | 4      | 3,2  |
|                                                               | Stellenbosch Local Municipality           | 2      | 1,6  |
| City of Cape Town<br>Metropolitan<br>Municipality (47 =37.9%) | Cape Town Eastern Health sub-District     | 14     | 11,3 |
|                                                               | Cape Town Northern Health sub-District    | 6      | 4,8  |
|                                                               | Cape Town Southern Health sub-District    | 1      | 0,8  |
|                                                               | Khayelitsha Health sub-District           | 12     | 9,7  |
|                                                               | Tygerberg Health sub-District             | 14     | 11,3 |
| Overberg District                                             | Overstrand Local Municipality             | 2      | 1,6  |
| Municipality (3=2.4%)                                         | Theewaterskloof Local Municipality        | 1      | 0,8  |
| West Coast District<br>Municipality (3=2.4%)                  | Bergrivier Local Municipality             | 1      | 0,8  |
|                                                               | Matzikama Local Municipality              | 1      | 0,8  |
|                                                               | West Coast District Managed Area          | 1      | 0,8  |

## INFRASTRUCTURE DEVELOPMENT

## **Tygerberg Children's Hospital Trust Contribution**

## MEDICAL EQUIPMENT PROCURED FOR THE PERIOD

| Date       | Payment made to:                      | For:                                                              | Amount       |
|------------|---------------------------------------|-------------------------------------------------------------------|--------------|
| 25/01/2018 | East Coast Medical                    | Cabinets & storage carts                                          | R 206 872,24 |
| 14/02/2018 | Grobir Medical<br>Suppliers (Pty) Ltd | 3 X Oxygen saturation monitors for<br>Bronchoscopy & G9           | R 67 857,76  |
| 26/02/2018 | East Coast Medical                    | 1 Storage Transportation Cart for Paediatric Oncology             | R 30 181,67  |
| 30/07/2018 | East Coast Medical                    | Paediatric Oncology (6 mini carts + 1 storage transportation cart | R 72 396,95  |
| 05/09/2018 | Tecmed Africa                         | Toshiba probe for transcranial doppler - Oncology                 | R 122 393,73 |
| 12/12/2019 | Medsci (Pty) Ltd                      | Inbody S10 with accessories                                       | R 173 164,00 |
| Total      |                                       |                                                                   | R 672 866,35 |

## **UPGRADE OF FACILITIES FOR THE PERIOD**

|            |                                      | •                                                               |              |
|------------|--------------------------------------|-----------------------------------------------------------------|--------------|
| Date       | Payment made to:                     | For:                                                            | Amount       |
| 06/07/2018 | De Klerk Painters & Maintenance      | Painting of Ward A9                                             | R 12 750,00  |
| 05/09/2018 | AC Flooring Services CC              | Flooring Ward A9                                                | R 25 190,70  |
| 05/09/2018 | De Klerk Painters & Maintenance      | 50% deposit - painting Ward A9                                  | R 15 930,00  |
| 03/10/2018 | De Klerk Painters & Maintenance      | Ward A9 - painting & fixing of corridor                         | R 16 550,00  |
| 29/10/2018 | De Klerk Painters & Maintenance      | Maintenance to Ward A9                                          | R 20 000,00  |
| 29/10/2018 | Dustin Juries                        | Kitchen cupboards Ward A9                                       | R 9 500,00   |
| 12/11/2018 | De Klerk Painters and<br>Maintenance | Ward A9 - painting of 43 doors                                  | R 8 800,00   |
| 19/11/2018 | Dustin Juries - JTD's                | Ward A9 - kitchen cupboards (bottom)                            | R 4 800,00   |
| 21/11/2018 | Jaco De Klerk                        | Billing for additional work done to corridor of Ward A9         | R 9 300,00   |
| 06/12/2018 | De Klerk Painters & Maintenance      | 50% balance payment for painting and maintenance A9             | R 15 930,00  |
| 12/12/2018 | De Klerk Painters & Maintenance      | Ward A9 - preparation for painting corridor walls & door frames | R 54 650,00  |
| Total      |                                      |                                                                 | R 193 400,70 |

## DONATIONS RECEIVED FOR THE PERIOD

| DATE                     | SUPPLIER                                 | PROJECT/WARD                                                     | AMOUNT              |
|--------------------------|------------------------------------------|------------------------------------------------------------------|---------------------|
| 05/01/2018               | Roskim's Christmas<br>Lights             | A9 ICU project                                                   | R 7 908,20          |
| 08/01/2018               | Roskim's Christmas<br>Lights             | A9 ICU project                                                   | R 250,00            |
| 19/01/2018<br>19/01/2018 | Sanbonani Holdings<br>Sanbonani Holdings | Kangaroo Chairs Epoxy coated X3 Bedside Lockers Cream X3         | R14 550,00          |
| 31/01/2018               | Bosmansdam High –<br>Donation            | Paediatric Oncology                                              | R 6 000,00          |
| 22/02/2018               | Dis-chem Pharmacies                      |                                                                  | R449,00<br>R1179,60 |
| 23/03/2018               | Sanbonani Holdings                       | Children's cotX3                                                 |                     |
| 25/03/2018               | Ms Logandree Gounden                     | Paediatric Oncology                                              | R 10 000,00         |
| 19/04/2018               | Mrs B. Lotz                              | Television Set Hisense Flat screen                               | R3000,00            |
|                          | Mrs B. Lotz                              | Television Set Diamond 24 Inch                                   | R3000,00            |
|                          | Mrs B. Lotz                              | Television Set Sinotec 24 Inch                                   | R3000,00            |
|                          | Mrs. B. Lotz                             | Television Set Sinotec 32 Inch                                   | R3000,00            |
|                          | Mrs. B. Lotz                             | Television Set Diamond 32 Inch                                   | R3000,00            |
| 24/04/2018               | Sanbonani Holdings                       | Movable table X3                                                 | R8 472,84           |
| 30/07/2018               | Oasis Group                              | A9 UCI project                                                   | R 10 000,00         |
| 07/09/2018               | Oasis Group                              | Purchase a transcranial Doppler probe for<br>Paediatric Oncology | R250 000,00         |
| 07/12/018                | Accordian Street Primary School          | Paediatric Oncology                                              | R1000,00            |
| 31/12/2018               | Roskim's Christmas<br>Lights             | A9 ICU project                                                   | R31 880,00          |
| 08/2018                  | Dis-chem<br>Pharmacies                   |                                                                  | R7417.65            |
| 27/09/2018               | Sanbonani Holdings                       | Kangaroo Chair Epoxy and Plastic Coated X1                       | R4 850.00           |
| Total                    |                                          |                                                                  | R332 139,64         |

## Part 2

## Community Outreach Programmes/Community Service and Interaction & Partnerships

## **Expert Members**

#### Prof M Kruger

- Member of the Central Clinical Committee of the South African Health Products Regulatory Authority (SAHPRA) since November 2014
- Member of the Provincial Clinical Governance Committee (PCGC)
- Member of the Senate Research Ethics Committee of Stellenbosch University
- Member of the Research Ethics Executive Committee of the Faculty of Medicine and Health Sciences, Stellenbosch University
- Member of the Postgraduate Research Committee of the Faculty of Medicine and Health Sciences, Stellenbosch University
- Chair of the Health Research Ethics Committee 1 of the Faculty of Medicine and Health Sciences, Stellenbosch University
- Trustee of the Tygerberg Children's Hospital Trust

## **Prof S Kling**

- Immediate Past President, Council of College of Paediatricians
- Ethics accreditor, CPD Committee, Faculty of Medicine and Heath Sciences.
- Vice-chair, Undergraduate Research Ethics Committee since 2015
- Member of the Research Ethics Executive Committee of the Faculty of Medicine and Health Sciences, Stellenbosch University
- Member, Clinical Ethics Committee Tygerberg Hospital (2002 2004 and 2009 to current).
- Allergy Society of South Africa (ALLSA)
  - Member of Executive Committee since 1998
  - o Chairman, Research Subcommittee (2011 to date)

#### **Prof AC Hesseling**

- NRF SARChi Chair in Paediatric Tuberculosis.
- Associate editor: International Journal against Tuberculosis and Lung Diseases, since 2009
- Site PI, CDC TBTC Network, Site 33, Stellenbosch University, since 2009
- Chair, Tuberculosis Scientific Committee, IMPAACT network, since 2011
- Co-chair: Pediatric Interest Group, TBTC network, since 2010
- Consultant: TB Alliance, Pediatric TB Unitaid program, since 2013
- Co-chair: PhD Committee, Department of Paediatrics and Child Health, Stellenbosch University, since 2015
- Member, Research committee (subcommittee C), Faculty of Medicine and Health Sciences, Stellenbosch University, since 2016
- South African TB Think Tank, TB Prevention Task force core member, since 2017
- Consortium Chair: South African Gates TB Demonstration and Scale-up, since 2018

#### **Prof MF Cotton**

- Member of the Clinical Guideline Development Group (GDG) to support the update of the World Health Organization Consolidated Guidelines on the use of antiretroviral drugs for treating and preventing HIV infection
- WHO HIV/TB Task Force Advisor

## Prof N Beyers

POPART Zambia & RSA

#### Prof HS Schaaf

- Board of Directors and Chairperson of Adult and Child Lung Health section, The Union (International Union Against Tuberculosis and Lung Disease)
- Expert on ATS/CDC committee for new TB guidelines
- MDR-TB Review Board Western Cape
- Member of the Subcommittee C of the Research Committee, Faculty of Medicine and Health Sciences

#### Dr B Fourie

 Safety representative on Health and Safety Committee, Faculty of Medicine and Health Sciences

#### Prof P Goussard

- Health Research Ethics Committee, Faculty of Medicine and Health Sciences
- Chair of the Tygerberg Children's Hospital Trust

#### Prof A Bekker

- International Maternal Pediatric Adolescent AIDS Clinical Trials Network (IMPAACT) P1106 vice-chair
- CDC: Tuberculosis Trials Consortium (TBTC) Tuberculosis and Pregnancy Working Group membership
- Health Research Ethics Committee, Faculty of Medicine and Health Sciences
- Paediatric HIV advisory Board for ViiV
- Collaborating with GARDP/DNDi on a NeoAMR global neonatal sepsis study

#### Prof J Lawrenson

• Ethics and Guidelines committee – SA Heart Association

#### Dr S Holgate

USANA executive committee member – treasurer

## Dr G Kali

 Neonatal Encephalopathy with Suspected Hypoxic Ischemic Encephalopathy (NESHIE) study local PI

#### Prof R van Toorn

• PANDA executive committee member –treasurer

### Dr CJ du Buisson

• South African Renal Society Executive Committee

## Outreach

### International

Profs PB Hesseling & M Kruger – Childhood cancer Cameroon

#### **Regional and District**

- Worcester Profs R van Toorn, R Solomons, J Lawrenson, B Fourie & Dr C du Buisson
- Paarl Profs P Goussard, H Rabie, R van Toorn, R Solomons, Lawrenson and Dr B Fourie. HIV outreach: Dr Frigati/Rose
- George Prof J Lawrenson, Dr B Fourie
- East London Prof J Lawrenson, Dr B Fourie
- Delft Drs L Smit, R Gioio (HIV outreach)
- Helderberg Dr T Wessels; Perinatal Dr H Hassan, Dr Frigati/Rose, (HIV outreach)
- Khayelitsha district hospital Prof H Rabie, Dr P Rose
- Eersterivier Drs H Finlayson (HIV outreach), JCF du Preez

- Ikwezi clinic HIV outreach
- Bishop Lavis clinic, HIV outreach
- Grabouw CHC HIV outreach
- Kraaifontein CHC HIV outreach
- Scottsdene MDR TB outreach clinic Prof Schaaf, Dr Rose
- Town 2 Khayelitsha MDR TB outreach clinic Prof Schaaf, Dr Rose
- City of Cape Town DTTC
- DTTC skills development in the community
- Karl Bremer Hospital, Haemophilia, Childhood Cancer Dr A van Zyl; Perinatal Dr H Hassan, Prof S Kling (General Paediatrics) and Dr H Finlayson (Antibiotic Stewardship), Neonatal – Neonatal team
- George Dr B Fourie, Prof J Lawrenson
- East London Prof J Lawrenson
- Brooklyn Chest Hospital Prof HS Schaaf, Dr H Finlayson
- Paarl School for Children with Cerebral Palsy, Brackenfell---Dr Cilla Springer
- Western Cape Education Department outreach meeting for autism spectrum disorder, Chere Botha Centre, Bellville – Dr Cilla Springer
- Neurodevelopmental clinic HOPE doctors and occupational therapist have been assisting once a week in a voluntary basis.

## **Tygerberg Children's Hospital Trust**

## Trustees: Prof P Goussard (Chair), Dr D Erasmus, Profs M Kruger, J Smith

The TCH Trust had a number of successful events in 2018, including the annual golf day. In 2018 the trust started a reconstruction process, which will continue into 2019. The plan is a major overhaul of the current trust, including new trustees and also the employment of a company to run the daily administrative of the trust and to launch some major fund-raising events. A number of projects have been started and we will continue to be developed in the coming years. We also had radio campaigns during the early part of 2019 in efforts to create awareness of the trust under local communities. Snapscan accounts have been created for easier donation of funds. New systems of Champions of the trust have been created. In 2018 the trust also was selected as one of a few, being officially supported by the Arch Bishop Tutu Foundation. We would also like to thank everyone who supported the trust in 2018 and for their continuing support in 2019. We had many donations in 2018, not only financial support but also the donation of medical equipment.

We value all of these contributions as they not only improve the medical care of the patients but also improve the comfort of the mother and child during their hospital stay.

## **HOPE Cape Town**

## Drs I Reyneke, M Esser

HOPE Cape Town is an NGO (Rotary Signal Hill and the German Speaking Catholic Community Cape Town) with offices in the department and link community paediatric HIV and TB community care with the department. In ward G7 HOPE CT has re-established a playroom, well stocked with age appropriate toys – specifically selected by the HOPE Occupational Therapist. Volunteers are involved to entertain and engage with children three times per week. A crafts program has been established in G7 in 2018 with a facilitator who visits the ward twice a week – teaching caregivers various craft techniques – knitting, crocheting, needle felting and. Caregivers also receive a support "starter pack" to keep them occupied during the long hours they spend next to their children's beds. HOPE CT employs two doctors both of whom are experienced HIV/ARV clinicians. These doctors assist the Tygerberg Infectious Diseases doctor to initiate and manage paediatric patients on ARVs at the Delft

CHC on a Friday. On average 26 children are seen per week at this clinic requiring comprehensive care. HOPE CT has also established an ART youth clinic Delft CHC – an adolescent friendly afternoon clinic – as well as adolescent support groups. For research support In 2018, HOPE CT co-funded Dr Lisa Frigati's study: Serologic correlates of protection against vaccine-preventable disease in Perinatally HIV-infected (PHIV) Adolescents at Tygerberg Hospital.

## Hear the TYGER ROAR (Reach out and Read) PROJECT at Tygerberg Hospital

#### Dr JCF du Preez

The Baby Book Project: We are aiming to move 16 000 books into patients homes this year. Currently, all children admitted or seen at Tygerberg Hospital receive a nursery rhyme book in their native language. The book includes a referral "script" to the closest library and book sharing tips for parents.

Volunteers service the outpatient's waiting rooms regularly. They hand out picture books from a book trolley and explain the importance, fun and joy of reading to the parents. Waiting rooms also have permanent bookshelves and children can browse through the books while waiting to be seen by the doctors. Medical students take the book trolleys around the wards once a week to ensure that in-patients share in the book reading project..

## Media Exposure

## TV/Radio Interviews:

 DSTV (KYKNet) - Die dag in die lewe van 'n ouer met 'n kind met kanker, 10 April 2018: Prof M Kruger

• Bush Radio interview: meningitis - 25 June 2018: Dr P Rose

#### **Articles:**

Name of publication: Tygerburger

Name of article: Generous donation made to Tygerberg (Profs M Kruger, P Goussard, J Smith, Dr N

Parker)

Date: 24 January 2018

Name of publication: VIVUS Medicine and Health Sciences

Name of article: SU researchers receive prestigious Fogarty awards (Prof A Dramowski, Dr A

Slogrove)

Date: March 2018

Name of publication: Stellenbosch University Web News

Name of article: 'Om gemeenskap te help' die motivering after Rory se PhD (Dr R Dunbar)

Date: 20 March 2018

Name of publication: Stellenbosch University Web News

Name of article: SU researchers receive prestigious Fogarty awards (Prof A Dramowski, Dr A

Slogrove)

**Date**: 17 April 2018

Name of publication: Stellenbosch University Web News

Name of article: Ukwanda bied kritieke navorsing platform (Prof N Beyers)

Date: 5 Junie 2018

Name of publication: Timeslive

Name of article: Desmond and Leah Tutu spend wedding anniversary with team caring for sick

children (Profs M Kruger, N Beyers, RP Gie, Dr A van Zyl)

**Date**: 2 July 2018

Name of publication: News24

Name of article: Archbishop Tutu and wife spend wedding anniversary at Tygerberg Hospital

**Date**: 2 July 2018

Name of publication: Stellenbosch University Web News

Name of article: FMHS researchers reveal pivotal work on managing TB, HIV (Profs M Kruger, N

Beyers, Drs P Bock, F Marx, S-A Meehan, R Dunbar

**Date**: 10 April 2018

## **International visitors:**

**Prof Paul Rogers**, Clinical Professor, University of British Columbia & BC Children's Hospital, 10 April 2018:

- Provided expert advice as co-supervisor towards the PhD of Dr Anel van Zyl titled: 'Long-term follow-up and care of childhood cancer survivors in South Africa' and Ms Judy Schoeman titled: 'Nutritional interventions in children with cancer.' (both are supervised by Prof Mariana Kruger)
- Led training rounds in Tygerberg Hospital for senior registrars in Paediatric Oncology
- Discussed future collaboration between Stellenbosch University and University of British Columbia

**Archbishop Desmond Tutu and Mrs Leah Tutu** celebrated their wedding anniversary at Tygerberg Children's Hospital, 2 July 2018.

**Prof Einspieler,** University of Austria, General movement trust. 22 – 26 August 2018

**Prof Chris van Geet**, Vice Rector of Biomedical Sciences Group, Katolieke Universiteit, Leuven - Platelet function disorders, 28 August 2018

**Prof Mirjam M van Weissenbruch**, Pediatrician-Neonatologist, Department Pediatrics /IC Neonatology, Amsterdam UMC - The vulnerable new-born, how feasible is a healthy later life? – 19 September 2018

**Prof Monika Bullinger** gave a lecture on "Quality of life in children with chronic conditions", on 3 December 2018.

**Prof Mary Rutherford;** Neuroradiologist, Kings College London. Neonatal Brain MRI discussions as co-supervisor of Dr Gugu Kali's PhD.

#### Part 3

## **Teaching & Training**

## **Education-related Activities**

## **Postgraduate Students**

## **Successful PhD Candidates**

- **Dr P Bock:** Title: Impact of a Universal Test and Treat strategy on clinical outcomes amongst HIV infected adults in South Africa: Supervisor: Prof N Beyers, Dr S Fidler
- **Mr R Dunbar:** Title: How can virtual implementation modelling inform the scale-up of new molecular diagnostic tools for tuberculosis? Supervisor: Prof N Beyers, Mr I Langley
- **Dr F Marx**: Title: Mathematical modelling to project the impact of interventions targeted to previously treated individuals on the trajectory of the tuberculosis epidemic in high tuberculosis prevalence settings Supervisors: Prof N Beyers, Prof T Cohen
- **Dr S-A Meehan**: Title: The contribution of a community-based HIV counselling and testing (HCT) initiative in working towards increasing access to HIV counselling and testing in Cape Town, South Africa. Supervisors: Proff N Beyers, R Burger
- **Dr AJ Garcia-Prats**: Title: Addressing critical knowledge gaps to improve and shorten MDR-TB treatment regimens in children. Supervisors: Profs Anneke Hesseling and Simon Schaaf
- **Dr E Walters**: Title: Novel diagnostic strategies and markers of treatment response for paediatric pulmonary tuberculosis. Supervisors: Proff AC Hesseling, RP Gie

## **Enrolled PhD students**

- Mr G Mbah Afungchwi: Title: Advocacy for childhood cancer in Cameroon: improving early diagnosis and referrals to life-saving care. Supervisor: Prof M Kruger, Prof PB Hesseling
- Dr K du Preez: Title: Complementary surveillance strategies and interventions to inform a tuberculosis care cascade for children. Supervisors: Proff AC Hesseling, HS Schaaf and Dr P Naidoo
- **Dr GTJ Kali**: Title: A comparative study of neuroprotective strategies in neonatal hypoxic ischaemic encephalopathy. Supervisors: Prof J Smith, Dr M Rutherford
- **Dr Y Kooblal:** Title: The relationship of body composition on nutritional and immunological status in childhood tuberculosis. Supervisors: Prof Ronald van Toorn, Dr Martijn van der Kuip and prof Marceline Tutu van Furth
- **Dr B Laughton:** Title: The effects of early versus delayed antiretroviral treatment on the short- and long-term neurodevelopmental outcome of children who are HIV positive. Supervisor: Prof MF Cotton, Prof M Kruger
- **Dr M Osman:** Title: TB-associated mortality in South Africa: longitudinal trends and the impact of health system interventions. Prof AC Hesseling and Dr P Naidoo
- Prof L Pepeta: Title: Novel medical and percutaneous modalities in the management of a
  patent ductus arteriosus in middle income and low income countries. Supervisors: Prof M
  Kruger, Dr HN Ntsinjana
- **Prof H Rabie**: Title: Pharmacokinetics and therapeutic outcomes of children with tuberculosis/HIV co-infection treated with lopinavir/ritonavir and a rifampicin-containing anti-tuberculosis regimen. Supervisors: Proff MF Cotton, HS Schaaf, RP Gie

- **Me J Schoeman:** Title: Nutritional status of children at cancer diagnosis and during treatment, with a focus on the association with their clinical outcome. Supervisors: Profs M Kruger, Elena J. Ladas and Paul Rogers.
- **Dr P Springer:** Neurodevelopmental Outcome of the HIV exposed but uninfected Infant and evaluation of a developmental screening tool. Supervisors: Prof Mariana Kruger, Prof Christopher Molteno
- **Dr J van Heerden**: Title: Neuroblastoma In Low And Middle Income Countries. Supervisors: Prof M Kruger
- **Dr L van Wyk:** Title: Non-invasive cardiac output monitoring in preterm infants . Supervisors: Prof Johan Smith, Dr John Lawrenson and Prof Willem-Pieter de Boeck
- **Dr A van Zyl:** Title: Long-term follow-up and care of childhood cancer survivors in South Africa. : Supervisors: Profs M Kruger and P Rogers

## Successful MMed(Paed) Candidates

- Dr E du Plooy
- Dr B Wentzel
- Dr S Abumhara
- Dr I Abrahams
- Dr C Richardson
- Dr L Scheepers
- Dr L Schoonraad
- Dr K Oppel
- Dr L van Heerden
- Dr L Mfingwana
- Dr F Adenivi
- Dr SH Fry

## **Colleges of Medicine of South Africa (CMSA)**

- Certificate in Paediatric Pulmonology
  - o Drs Lindy-Lee Green with distinction (Connor Farrel Medal)
- Certificate in Medical Oncology
  - o Dr Lillian Gesami-Steytler
- Certificate in Neonatology
  - o Drs Natalie Paulse, Mary Kariuki, Onwona Agyeman
- FC Paed (SA) Part II
  - o Dr S Marais
  - o Dr F Adeniyi
  - o Dr L Mfingwana
  - o Dr L Schoonraad
  - o Dr L Scheepers
  - o Dr M van Velden
  - o Dr L van Heerden

- o Dr K Oppel
- o Dr CJ Richardson

## • FC Paed (SA) Primary

o Dr C van der Merwe (Leslie Rabinowitz Medal)

## **University of Stellenbosch Business School (USB)**

Dr I Appel - Postgraduate Diploma in Business Management and Administration (PGD BMA)

## **Undergraduate Students**

98% pass rate (2017 = 99%)

## **Education-related Activities**

- The Education Committees of the Department comprised as follows:
  - o Undergraduate: Dr L Smit (Chairperson), Dr A Redfern (secundus)
    - MBChB II: Drs P Rose, R Uys, E Malek, S Holgate, L LLoyd
    - Early rotation: Drs I Appel, L Frigati, prof H Rabie, H Hassan, Drs A Thomas, M Morkel, Sr L Yzelle
    - Middle rotation: Drs N Parker, B Fourie, C du Buisson, H Finlayson, G Kali, A Bekker
    - SI: Prof R Solomons, Drs JCF du Preez, L Smit, A van Zyl, S Holgate, C Springer, Prof E Zöllner, Drs M Morkel
    - Remedial: Dr M Morkel, Sr L Yzelle
    - Elective students: Drs H Finlayson, I Appel, Prof M Kruger
  - Postgraduate: Dr G Kali (chairperson), Proff R van Toorn, A Bekker, M Kruger, ED
     Nel, H Rabie, P Goussard, HS Schaaf
  - o PhD: Proff M Kruger (Chairperson), MF Cotton, AC Hesseling, HS Schaaf, J Smith
- All consultants are involved with under- and postgraduate teaching on a daily basis.
   Additional education activities included:
- External examiners at universities in South Africa:
  - Undergraduate:
    - Dr JCF du Preez: Sefako Makgatho Health Sciences University
    - Drs Y Kooblal, N Paulse, L Smit: University of Kwazulu-Natal
- Senior registrars in training:
  - o Neonatology: Drs AD Daniels, N Paulse, A van Eck, I Abrahams, M Kariuki (Supernumerary from Kenya), K Onwona-Agyeman (Supernumerary from Ghana)
  - Paediatric Oncology: Drs L Schoonraad, L Gesami-Steytler (Supernumerary from Namibia)
  - o Paediatric Rheumatology: Dr DR Abraham
  - o Paediatric Pulmonology: Drs Y Mulambia, L-L Green
  - o Paediatric Neurology: Dr T Katangwe
  - Paediatric Gastroenterology: Drs LT Radebe, U Chowdhury, sponsored by a Nestle Nutrition Institute Fellowship endorsed by the European Society of Paediatric Gastroenterology
- Educational Outreach:
  - Prof M Kruger Accreditation visit (accreditor with Prof Vic Davies), CMSA College Paediatrics, Botswana, 5-6 March 2018

- Prof M Kruger Paediatrics and Child Health Research Day, UCT, 30-31 October 2018. Judge for best research with Prof Marion Jacobs.
- Colleges of Paediatricians (CMSA)
  - o Prof Ronald Van Toorn (Convenor) and Etienne Nel (co-convenor)
  - Prof Mariana Kruger examiner FCPII & Moderator, Cert Medical Oncology
  - Prof Sharon Kling Convenor, Cert Allergology (Fam Phys)(SA); examiner Dip Allerg(SA), Moderator FCPaed I & II
  - Dr Anel Van Zyl convenor, Cert Medical Oncology
  - o Dr. Ilse Appel observer Cert Critical Care
  - o Prof Regan Solomons examiner FCPII
  - Prof Johan Smith convenor Cert Neonatology
  - Prof Adrie Bekker- examiner Cert Neonatology
  - o Dr Sandi Holgate examiner Cert Neonatology
  - Dr Gugu Kali Examiner Cert Neonatology
  - o Dr Pricilla Springer convenor Cert in Neurodevelopment
  - o Prof Etienne Nel convenor GIT certificate exams
  - Dr Andrew Redfern examiner DCH exams
  - Dr Amy Slogrove examiner DCH exams
  - o Dr Thandi Wessels moderator DCH exams
  - Monika Esser convenor, Cert Rheumatology
  - Prof Pierre Goussard convenor Cert Pulmonology Diseases
  - Dr Christelle du Buisson convenor Cert Nephrology
  - Prof Rabie convenor Infectious diseases
  - Dr Barend Fourie, convenor, Cert Cardiology
  - Prof Lawrenson examiner , Cert Cardiology
- South African Research Ethics Training Initiative (SARETI) UKZN, funded by Fogarty International Fogarty International Center (FIC) of the National Institutes of Health (NIH), grant number 6R25TW001599-13: Prof M Kruger (Co-principal Director)

#### Part 4

## **Achievements and Highlights**

## **Appointments**

## **Prof Sharon Kling**

Re-elected onto the Executive Committee of the Allergy Society of South Africa.

## **Prof Pierre Goussard**

Elected to the South African Thoracic Society (SATS) Council.

**Prof Adrie Bekker, Dr Heather Finlayson and Dr Angela Dramowski** will serve on the newly formed National Neonatal Sepsis Task Force to advise the NDOH on surveillance, outbreak response, infection prevention and control, antimicrobial stewardship and case management of neonatal sepsis, focused particularly on antibiotic resistant infections and hospital-acquired sepsis in the public- and private-sectors.

## **Dr Amy Slogrove**

Appointed as a member of the South African Young Academy of Science.

## **Grants/Awards**

## **Prof ED Nel**

Awarded an Erasmus+ Capacity Building Grant: to Improve Early Nutrition and Health in South Africa (ImpENSA)

## Dr Lizelle van Wyk

The Southampton Faculty nominated her for the Deans List – which was awarded in October 2018 for achieving the highest academic performance in the year and being nominated by a tutor performance or achievement.

She is doing her MSc Neonataology with an MSc by dissertation, an on-line degree from the University of Southampton, United Kingdom. The final dissertation was a systematic review of the accuracy and precision of non-invasive cardiac output monitoring methods in neonates with a PDA.

## Dr Angela Dramowski

Awarded a Discovery Foundation Distinguished Visitor Award to work with Dora Nginza Hospital in the Eastern Cape over the next 2 years to improve Infection Prevention and Antibiotic Stewardship practices.

#### **Dr Barbara Laughton:**

Awarded NIH grant as part of the study: "The Association of Perinatal HIV infection and hearing loss in children of Cape Town, South Africa" with UCLA for 5 years.

#### **Prof Ekkehard Zöllner:**

Awarded NRF grant as part of the "Competitive Programme for Rated Researchers" for 3 years for project "Genetic predictors of HPA suppression in children treated with corticosteroids".

## NRF ratings:

| 0 | Prof PR Donald    | A2 |
|---|-------------------|----|
| 0 | Prof MF Cotton    | A2 |
| 0 | Prof HS Schaaf    | A2 |
| 0 | Prof AC Hesseling | B1 |
| 0 | Prof J Smith      | C2 |
| 0 | Prof R Van Toorn  | C2 |
| 0 | Prof P Goussard   | C2 |
| 0 | Prof PB Hesseling | C2 |
| 0 | Dr A Dramowski    | C3 |
| 0 | Prof EWA Zöllner  | C3 |
| 0 | Dr A Bekker       | C3 |
| 0 | Dr H Rabie        | C3 |
| 0 | Dr S Innes        | Y2 |
| 0 | Prof R Solomons   | Y2 |

## **Sabbaticals**

## **Prof Sharon Kling** - 1-28 February 2018

Prof Kling is enrolled at Stellenbosch University for her PhD in Applied Ethics, titled: 'Models of clinical ethics consultation services in South Africa: A critical appraisal.' During her sabbatical leave Prof Kling wrote up 4 chapters of her thesis.

#### Prof Adrie Bekker - 28 May to 27 July 2018

Prof Bekker was the recipient of The Collaborative Initiative for Paediatric HIV Education and Research (CIPHER) Growing the Leaders of Tomorrow Fellowship Programme of the International AIDS Society (IAS). Funding of a total amount of US \$70,000.00 was awarded to her for the research project entitled "Pharmacokinetic and safety characteristics of nevirapine and lopinavir/ritonavir in HIV-exposed and infected low birth weight infants".

Prof Bekker attended pharmacology training in San Diego, the IMPAACT annual meeting in Washington and the International AIDS conference in Amsterdam.

Dr Elmarie Malek - 15 October to 14 December 2018

During her sabbatical, Dr Malek developed projects for the Provincial Department of Health:

- Project 1: Document the process and outputs of the PSG3 Parent Infant Child Health and Wellness Working Group May 2013- October 2018
- Project 2: Produce a 1st draft of a Framework for a Provincial DOH Well Child Policy

## **Organizing and Presentations at Conferences**

The **seventh annual Here by Lungs Congress,** organised by Profs P Goussard and J Smith, was held on 15-16 March 2018 at Spier, Stellenbosch.

#### Goussard, P

Bronchoscopy in PTB, 14 March 2018

Interventional Pulmonology, 15 March 2018

The department successfully organised the **bi-annual ALLPAEDS conference** (the joint South African Paediatric Association (SAPA) & Allergy Society of South Africa (ALLSA) conference), held at Century City Conference Centre from 29 August – 2 September 2018.

Presentations were also given by members of the department as follows:

## **Invited Speakers:**

- Dr Ilse Appel Adverse events
- Dr Angela Dramowski IPC management of drug resistant bacterial organisms in the hospital
- Prof Anneke Hesseling TB exposure in the preschool setting
- Dr Barend Fourie
  - Paediatric cardiology for the general paediatrician 2018: the role of ECG in the care of the patient
  - Paediatric cardiology for the paediatrician 2018: the role of CXR interpretation in the care of the patient
- Prof Pierre Goussard Imaging of Pneumonia
- Prof Etienne Nel
  - Functional gastrointestinal disease: how can the general paediatrician use the Rome IV criteria?
  - o Recognising IBD in children
- Prof Sharon Kling Futility at end-of-life care who decides? (Ethics)
- Prof Mariana Kruger Emergencies in Paediatric Oncology
- Prof John Lawrenson Paediatric cardiology for the paediatrician 2018: preventative cardiology and child health
- Dr Julie Morrison Approach to persistent bacterial bronchitis
- Dr Andrew Redfern Investigating the paediatric patient with first time seizure
- Dr Penelope Rose Approach to chronic abdominal pain in children (GIT workshop at SAPA)
- Dr Lizelle van Wyk Minimally invasive SRT use in neonates
- Dr Anel van Zyl Late effects of childhood cancer
- Prof Ekkehard Zöllner Hyperglycaemic emergencies-pitfalls & challenges

## Selected abstracts for presentation:

• Dr Ilhaam Abrahams - The correlation between the placental pathology and the neurological outcome of neonates ≥ 36 weeks with neonatal encephalopathy treated with therapeutic hypothermia.

- Dr Elri Du Plooy PMTCT and clinical profile of hospitalized HIV-infected children in an era of high pMTCT uptake and efficacy (**Dr Du Plooy won the prize for the best registrar presentation**).
- Dr Ruth Arayo Clinical presentation and treatment outcomes of infants with tuberculosis in Cape Town, South Africa
- Dr Sebastian Gericke Changes in cerebral blood flow and cardiac output in premature infants in the first 72 hours of life
- Dr Barbara Laughton Neurodevelopmental outcome at 11 months in HIV-infected infants initiating very early Antiretroviral therapy
- Dr Barbara Laughton HIV infected children show decreased activation in the auditory cortex
- Dr Lydia Mapala Burden, pathogen spectrum and outcome of paediatric community-acquired infections at Tygerberg Hospital
- Dr Lyn Scheepers Assessing the rate and factors associated with unscheduled return visits to a medical paediatric emergency department in a resource limited setting
- Dr Leilah Schoonraad Descriptive review of the health supervision received by children with Down syndrome at Worcester Provincial Hospital from 2010 to 2015

## **Sessions Chairs:**

- Dr Ilse Appel SAPA SESSION CRITICAL CARE
- Prof Mark Cotton SAPA SESSION INFECTIOUS DISEASES
- Dr Christel du Buisson SAPA SESSION RENAL
- Dr Christel du Buisson & Prof Etienne Nel SAPA SESSION ABSTRACT PRESENTATIONS
- Prof Pierre Goussard THE CAS MOTALA JOINT PLENARY SAPA/ALLSA
- Dr Sandi Holgate USANA SESSION
- Prof Sharon Kling JOINT PLENARY SESSION 6 AND CLOSING SAPA/ALLSA
- Prof Mariana Kruger JOINT PLENARY SESSION 4 SAPA/ALLSA & SAPA SESSION ABSTRACT PRESENTATIONS
- Prof John Lawrenson SAPA SESSION CARDIOLOGY
- Prof Etienne Nel SAPA SESSION GASTROENTEROLOGY
- Dr Andrew Redfern GENERAL PAEDIATRICS
- Prof Regan Solomons PLENARY SESSION 2 SAPA
- Prof Ronald van Toorn PANDA
- Dr Anel van Zyl SAPA SESSION HAEMATOLOGY AND ONCOLOGY
- Prof Ekkehard Zöllner SAPA SESSION ENDOCRINOLOGY & PLENARY SESSION 5 SAPA

The **fourth Annual Paediatric Registrar Research Day** was held on Wednesday 7 November 2018 here in the Faculty where 6 registrars presented their research. The prize for best research was awarded to Drs Sebastian Gericke & Folasade Adeniyi.

The eight annual Paediatric Academic Day of Excellence (WOW! Women of Worth) was held on Wednesday 7 November 2018 in the Faculty. It included invited speakers: Dr Aneesa Vanker (UCT) and Prof Mphele Mulaudzi (Wits).

#### **OTHER CONFERENCES**

## Afungchwi, Mbah, Glenn

 Development of a competency-cased orientation programme for pediatric oncology nursing in rural Cameroon & Strategies and priorities for dissemination of the SIOP baseline nursing standards: leadership and advocacy workshop outcomes. SIOP 2018 Congress, Kyoto, Japan, 13-19 November 2018

#### Bekker, A

 Stewardship & Managing neonatal infection outbreaks. Kenyan Paed Assoc Annual Conference, Mombassa, Kenya, April 2018

#### Cotton, MF

- Pharmacokinetics, Safety, And Efficacy Of Bictegravir/Emtricitabine/Tenofovir Alafenamide (B/F/TAF) Single-tablet Regimen In Children (6 To <12 Years) Living With HIV-1 - International AIDS Conference, Amsterdam, 23-27 July 2018
- Organizing Committee 10<sup>th</sup> Paeds HIV Workshop, Amsterdam, 20 21 July 2018
- Organizing Committee , SA HIV Clin SOC Conference, JHB, 24 -27 October 2018

#### Donald, PR

 Paediatric tuberculosis and the pharmacokinetics/pharmacodynamics sciences: what do we know and what don't we know?; What is the right dose of isoniazid? 49th Union World Conference on Lung Health, The Hague, Netherlands, 24-27 October 2018:

## Goussard, P

- Imaging in complicated pulmonary TB & Neonatal IND. Pan African Thorasic Society/South African Thorasic Society Congress, Durban, 12 15 April 2018
- Roles and pitfalls of computed tomography and magnetic resonance imaging in intrathoracic TB in HIV+/-children. The Union Congress .The Hague Holland October 2018
- Managing foreign bodies of the airway. 54th South African ENT Congress and 4th World Congress on Cochlear Implants in Emerging Countries, Cape Town, 28 October – 2 November 2018

## Grobbelaar, M

• Lumbar cerebrospinal fluid evolution in childhood tuberculous meningitis. 14<sup>th</sup> International Conference on Clinical Pediatrics, 14-16 June, Rome, Italy 2018. (*Dr Grobbelaar received a certificate of recognition for her phenominal and worthy oral presentation*).

#### Kali, GTJ

• Invited speaker: *HIE – can we do better?* Kenyan Paediatrics Association Annual Conference, Mombassa, Kenya, April 2018

#### Kling, S

- Ethics of Paediatric Palliative Care. PG Dip in general and community at Red Cross, UCT, 01 March 2018
- Clinician integrity, patient autonomy and the patient's best interests; and Ethical dilemmas in clinical practice. GP refresher Course, 25 May 2018

## Kruger, M

- Population Specific Issues in HIV Cure Research: Implication for Children ETHICS. Regulation of clinical research related to HIV care Conference, Bethesda, MD, 23-26 January 2018.
- African Health Research Ethics Symposium III (AHRES): SARETI, Umhlanga Rocks, South Africa, 29-31 January 2018
- Opening Address. Here be Lungs 7 Conference, Spier Estate, Stellenbosch, 15-16 March 2018
- Ethical Issues on HIV Cure Research in Pediatric Populations. NIAID DAIDS workshop: "Framework for Children, HIV Cure Research & Ethics. Initiating Pediatric Studies of HIV Cure Interventions: Scientific Knowledge Gaps, Regulatory and Ethical Considerations", 21 – 25 May 2018, Rockville, MD.
- Ethical issues in paediatric clinical trials in both developed and developing countries. Improving the lives of patients with CF in the 21st century. Catholic University of Leuven Campus Gasthuisberg, UZ Leuven, 21-22 September 2018.
- Report of first national collaborative treatment protocol of Retinoblastoma in South Africa An interim analysis. 50th SIOP Congress of the International Society of Paediatric Oncology

(SIOP): November 16-19, 2018, Kyoto, Japan.

#### Nel, ED

SAGES Conference, Session Chair, Speaker: Paediatric Reflux Disease (GP Update),
 Johannesburg, 8-11Augustust 2018

#### Schaaf, S

- New/Repurposed Drugs/MDR TB Drug and Treatment in Children. 22nd Annual Conference of the Union-North America, Chicago, IL, USA.28 Feb 3 March 2018.
- 49<sup>th</sup> Union World Conference on Lung Health, The Hague, Netherlands, 22-26 October 2018:
  - Speaker: Introductions and overview of course
  - o Precentative treatment consideration sof multidrug-resistant tuberculsos
  - Treatment of multidrug-resistant tuberculosis: paediatrics
  - EP04-132-26 Culture-confirmed tuberculosis in infants less than 3 months of age: clinical presentation and management
  - Injectable-free regimens, and the role of injectable agents, in the treatment of children with DR-TB
  - Chair: PGC02 TB prevention in children in high TB burden settings: implementing the child contact management (CCM) care cascade (Postgradaute Course)

#### Slogrove, A

 Debate: All infants should receive oral antiretroviral prophylaxis throughout breastfeeding -10th Paeds HIV Workshop, Amsterdam, 21 July 2018

#### Solomons, RS

 Plenary, Challenges in early diagnosis of tuberculous meningitis, International child neurology congress, Mumbai, 15-18 November 2018

#### Springer, P

 Launch meeting of the European Union-AIMS-2 Trials, which has included the Safe Passage Cohort in a prospective study on "Early bioMarkers of autism spectrum disorder and other neurodevelopmental disorders", EU-AIMS, Munich, Germany, 13-15 June 2018

## **Conferences Attended**

#### Bekker, A

- Conference on Retroviruses and Opportunistic infections, Washington, 13-16 February 2018
- Conference on Retroviruses and Opportunistic Infections (CROI), Boston, Massachusetts, 4-7
   March 2018

#### Cotton, MF

- Infectious Diseases Week (IDSA), San Francisco, US, 3-7 October 2018
- InterAmerica Congress of Paediatric Infectious Diseases, Mexico, 28-30 November 2018

#### Du Preez, JCF

- Ipokrates-Managing a neuroprotective NICU, Torino, Italy, 21-25 October 2018
- Panorama Annual Neonatal Seminar Panorama Hospital, 10 November 2018

#### Finlayson, H

- Antibiotic Resistance MAC, Pretoria, 09 May 2018
- ICAN Conference, Century City, CPT, 09-11 July 2018

## Fourie, B

- Cardiology Congress, Sun City, JHB, 11-12 October 2018
- CSI Africa, Egypt, 29-30 November 2018

#### Innes, S,

 Conference on Retroviruses and Opportunistic Infections (CROI), Boston, USA, 4-7 March 2018

- U.S.-South Africa Program for Collaborative Biomedical Research (NIH SA-MRC) Grantee meeting, Durban, SA, 11 June 2018
- International AIDS Conference, Amsterdam, 23-27 July 2018

### Kali, G

- Neonatal Neurology Conference, London, UK, May 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018

# Kruger, M

AMC Sub-Saharan Africa (SSA) Meeting, Dar Es Salaam, Tanzania, 10-13 July 2018

# Lloyd, L

- Here Be Lungs, Spier Estate, CPT, 15 -16 March 2018
- INAC Congress, Belgium, 21-26 June 2018

# Malek, E

- Priorities in Perinatal Care Conference, KZN, 14 March-16 March 2018
- Violence in Childhood-High level learning engagement Conference, George, 10 May 2018
- 16<sup>th</sup> WAIMH World Congress, Rome, 25 May 01 June 2018
- Safe Passage Symposium, CPT, 06 July 2018
- Breastfeeding Symposium, 03-10 August 2018
- FARR workshop, CTICC, 15 August 2018
- ECD Strategy Workshop, Lentegeur Hosp., CPT, 21 August 2018
- South African Academy of Family Physicians 21<sup>st</sup> National Conference, CPT, 24 August 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018

#### Morrison, J

- SATS/PATS Conference, Durban, 12-13 April 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018

#### Nel. E

Fellow Weekend (Gastroenterology Foundation of South Africa). Spier, Stellenbosch, 2-4
 February 2018

# Parker, N

CCSSA Congress Presenter Durban, UKZN, Durban, 18 July 2018

#### Redfern, A

- BadEM Conference, Greyton, CPT, 22 March 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 29-31 August 2018
- APLS, Red Cross, CPT, 15-16 November 2018

# Rose, P

- ESPGHAN Conference, 08-11 May 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 29 August 2018
- Annual International Child TB training Course, Goudini, 29 August-02 Sept 2018
- 49<sup>th</sup> Union World Conference on Lung Health, The Hague, Netherlands, 31 October 2018

#### Schaaf, HS

SATS/PATS, Conference, Durban, 12-13 April 2018

# Smit, L

- New RTHB National Orientation session, Ruslamere Hotel and Conference Centre workshop, Evertsdal, CPT, 26Jan 2018
- SAAHE Congress, Durban, 27-29 June 2018

#### Solomons, R

- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018
- Convenor- Paediatric Epilepsy Training (PET) Course, Red Cross, CPT, Convention Centre, CPT, 19 October 2018

# Springer, P

SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018

#### Van Zyl, A

SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018

#### Van Toorn, R

- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 30-31 August 2018
- Paediatric Epilepsy Training (PET) Course, Red Cross, CPT, 19 October 2018

# Van Wyk, L

- Attending Conference, Germany, 29 May-01 June 2018
- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 31 August2018

#### Zollner. E

- SAPA Allpaeds Congress, Century City Convention Centre, CPT, 31 August 2018
- 18th International Congress of Endocrinology in Cape Town, 1-4 December 2019

# Courses Organized and Presented

# Appel, I

APLS - instructor, Skills lab, Red Cross Children's Hospital, March and November 2018

#### Cotton, M

- Paediatric HIV Consultation, Bethesda, USA, 21-23 May 2018
- 10th Paeds HIV Workshop, Amsterdam, 20 July 2018

#### Du Buisson, CJ

Instructor, Saving Young Lives skills course, UCT, Cape Town, February 2018

#### Lawrenson, J

APLS Instructors Course, 24 -25 May 2018

# Schaaf, HS

Annual International Child TB training Course, Goudini, 10-13 Sept 2018

### Solomons, R

 Paediatric Epilepsy Training (PET) Course, Red Cross, CPT, Convention Centre, CPT19 October 2018

# **Courses Attended**

# Afaa, T

 Pre-Doctoral Programme in Health Sciences SU, ADA Health Sciences SU, 12 - 23 February 2018

# Afungchwi, Mbah, G

• ADA Summer School, EDP Programme, US, 08-13 January 2018

# Du Preez, JCF

 PredOctoberoral programme in Health Sciences, Stellenbosch University, 12-23 February 2018

# Finlayson, H

Annual International Child TB training Course, Goudini, 13 September 2018

#### Gesami-Steytler, L

• Hemofilie lesson, Red Cross, UCT, 12March 2018

# Hassan H

 PredOctoberoral programme in Health Sciences, Stellenbosch University, 12-23 February 2018.

# **Holgate S**

NIV Worksop, Pinelands, CPT, NIV Worksop, Pinelands, CPT, 18 July 2018

# Innes, S

- 10th HIV Pediatrics Workshop, Amsterdam, 20 July 2018
- 2nd Adolescent HIV workshop. Cape Town, SA, 10-11 October 2018
- 20th Comorbidities and Adverse Drug Reactions in HIV workshop, New York, USA, 13-14
   October 2018
- 2nd Microbiome in HIV workshop, Bethesda, USA, 16-17 October 2018

# Kali, GTJ

 Safe Passage Symposium, Faculty of Medicine & Health Sciences Faculty, Stellenbosch University, Tygerberg, 6 July 2018

# Kooblal, Y

• PREDAC, Goudini, CPT, US, 17-19 July 2018

# Lloyd, L

Medtronic Capnography Master Class, Germany, 8, 9 October 2018

# Malek, E

- Slide+Guide Content Workshop, CPT, 09 May 2018
- Antenatal Class Curriculum Expert Workshop, Claremont, CPT, 11 April 2018
- Provincial Inter-Departmental Committee for ECD, CPT, 24 May 2018
- Safe Passage Symposium, CPT, 06 July 2018
- RTHB Training, George, 12-13 July 2018
- Metro RTHB Training, 27 July 2018
- Breastfeeding Symposium, 03-10 August 2018
- Multidisciplinary district level allergy training program, RCWMCH, CPT, 06 August 2018
- NRTHB Training, Bellville Health Park, CPT, 08 August 2018
- RTHB Training, UWC, CPT, 10 August 2018
- RTHB Training, Paarl, 17 August 2018
- Child PIP Workshop, Paarl, 20 August 2018
- ECD Strategy Workshop, Lentegeur Hospital, CPT, 21 August 2018

# Schoeman, J

• ADA Summer School, EDP Programme, US, 08-13 January 2018

#### Schoonraad, L

• Paediatric Brain Tumour Workshop, Groote Schuur, CPT, 07 December 2018

# Smit, L

- MBChB Curriculum workshop, Evertsdal, CPT, 16 Feb 2018
- MBChB 2020 Curriculum Workshop, Feathers Lodge, Durbanville, CPT, 03 Dec2018

#### Springer, P

- Autism Diagnostic Observation Schedule, Child and Adolescent Unit, Child Psychiatry Red Cross Children's Hospital, 13-15 June 2018
- GCP Refresher, Stellenbosch University, July 2018
- Safe Passage Symposium, Faculty of Medicine & Health Sciences Faculty, Stellenbosch University, Tygerberg, 6 July 2018

# Uys, R

- Haemophilia with Inhibitors lecture by international expert, Prof Giangrande, Red Cross, UCT, 12 March 2018
- PREDAC, Goudini, CPT, 17-19 July 2018

### Van Zyl, A

- Haemophilia with Inhibitors lecture by international expert, Prof Giangrande, Red Cross, UCT, 12 March 2018
- Paediatric Brain Tumour Workshop, Groote Schuur, CPT, 03 December 2018

# **Publications**

# Journal Articles (subsidised)

- 1. ALLWOOD BW, GILLESPIE R, BATEMAN M, OLCKERS H, TABORDA-BARATA L, CALLIGARO G, VAN ZYL-SMIT RN, COOPER CB, BEYERS N, BATEMAN ED. Five-year follow-up of participants diagnosed with chronic airflow obstruction in a South African Burden of Obstructive Lung Disease (BOLD) survey. SAMJ South African Medical Journal 2018; 108(2):138-143.
- 2. ALLWOOD BW, MAARMAN GJ, KYRIAKAKIS CG, DOUBELL AF. Post-pulmonary tuberculosis complications in South Africa and a potential link with pulmonary hypertension: Premise for clinical and scientific investigations. *SAMJ South African Medical Journal* 2018; **108**(7):529.
- 3. APPEL I. Airway management in the critically ill patient. Medical Chronicle October 2018
- 4. ARAUJO DA SILVA AR, MARQUES AF, BISCAIA DI BIASE C, ZINGG W, DRAMOWSKI A, SHARLAND M. Interventions to prevent urinary catheter-associated infections in children and neonates: a systematic review. *Journal of Pediatric Urology* 2018; **14**(6):556.e1-e6.
- 5. BARNARD BW, DU PLESSIS A, GOUSSARD P, PITCHER RD. Infantile intrathoracic large airway obstruction in a setting with a high prevalence of tuberculosis/HIV. *Paediatrics and International Child Health* 2018; **38**(2):106-112.
- 6. BEYERS N, BURGER R, DRAPER HR, MEEHAN S, NAIDOO P, SLOOT R. Factors associated with linkage to HIV care and TB treatment at community-based HIV testing services in Cape Town, South Africa. *PLoS One* 2018; **13**(4):1-13.
- 7. BOCK PA, FATTI G, FORD N, JENNINGS K, KRUGER J, GUNST C, LOUIS F, GROBBELAAR CJ, SHANAUBE K, FLOYD S, GRIMWOOD A, HAYES R, AYLES H, FIDLER S, BEYERS N. Attrition when providing antiretroviral treatment at CD4 counts >500cells/μL at three government clinics included in the HPTN 071 (PopART) trial in South Africa. PLoS One 2018; 13(4):e0195127.
- 8. BOCK PA, JENNINGS K, VERMAAK R, COX HS, MEINTJES G, FATTI G, KRUGER J, DE AZEVEDO V, MASCHILLA LS, LOUIS F, GUNST C, GROBBELAAR CJ, DUNBAR R, BEYERS N, ET AL. Incidence of Tuberculosis among HIV-positive individuals initiating antiretroviral treatment at higher CD4 counts in the HPTN071 (PopART) Trial in South Africa. *Journal of Acquired Immune Deficiency Syndromes* 2018; **77**(1):93-101.
- 9. BOIVIN M, BARLOW-MOSHA L, CHERNOFF M, LAUGHTON B, ZIMMER B, JOYCE C, BWAKURA-DANGAREMBIZI M, RATSWANA M, ABRAHAMS N, FAIRLIE L, GOUS H, KAMTHUNZI P, MCCARTHY K, DU PLESSIS J, COTTON MF, ET AL . Neuropshychological performance in African children with HIV enrolled in a multisite antiretroviral clinical trial. *AIDS* 2018; **32**:189-204.
- 10. BOND V, NGWENYA F, THOMAS A, SIMUYABA M, HODDINOTT G, FIDLER S, HAYES R, AYLES H, SEELEY J. Spinning plates: livelihood mobility, household responsibility and anti-retroviral treatment in an urban Zambian community during the HPTN 071 (PopART) study. *Journal of the International AIDS Society* 2018; **21**(s4):e25117.
- 11. BRADLEY J, FLOYD S, PIWOWAR-MANNING E, LAEYENDECKER O, YOUNG A, MANDLA NF, BWALYA J, BOCK PA, FIDLER S, AYLES H, HAYES RJ. Sexually Transmitted Bedfellows: Exquisite Association Between HIV and Herpes Simplex Virus Type 2 in 21 Communities in Southern Africa in the HIV Prevention Trials Network 071 (PopART) Study. *Journal of Infectious Diseases* 2018; 218(3):443-452.
- 12. BYAMUNGU NL, DU PREEZ K, WALTERS E, NACHEGA JB, SCHAAF HS. Timing of HIV diagnosis in children with tuberculosis managed at a referral hospital in Cape Town, South Africa. *International Journal of Tuberculosis and Lung Disease* 2018; **22**(5):488-495.
- 13. CHABALA C, TURKOVA A, THOMASON MJ, WOBUDEYA E, HISSAR S, MAVE V, VAN DER ZALM MM, PALMER M, KAPASA M, BHAVANI PK, BALAJI S, RAICHUR PA, DEMERS A, HODDINOTT G, HESSELING AC, COTTON MF, ET AL. Shorter treatment for minimal tuberculosis (TB) in children (SHITE): a study protocol for a randomised controlled trial. *Trials* 2018; **19**:237.

- 14. CHERNOFF M, LAUGHTON B, RATSWANA M, FAMILIAR I, FAIRLIE L, VHEMBO T, KAMTHUNZI P, KABUGHO E, JOYCE C, ZIMMER B, ARIANSEN JL, JEAN-PHILIPPE P, BOIVIN M. Validity of Neuropsychological Testing in Young African Children Affected by HIV. *Journal of Pediatric Infectious Diseases* 2018; **13**(3):185-201.
- 15. COTTON MF, LIBERTY A, TORRES-ESCOBAR I, GONZALEZ-TOME MI, LISSENS J, ZARU L, KLAUCK I, CAMBILARGIU D, PIKORA C, CORRELL TA. Safety and efficacy of atazanavir powder and ritonavir in HIV-1-Infected infants and children from 3 months to <11 years of age. *Pediatric Infectious Disease Journal* 2018; **37**(6):e149-e156.
- 16. CRANMER LM, DRAPER HR, MANDALAKAS A, KIM S, MCSHERRY GD, KRZESINSKI EI, DU PLESSIS J, MITCHELL C, NACHMAN S, VAN DER LINDE MV, COTTON MF, HESSELING AC. High incidence of tuberculosis infection in HIV-exposed children exiting an isoniazid preventive therapy trial. *Pediatric Infectious Disease Journal* 2018; **37**(10):e254-e256.
- 17. CRICHTON H, O'CONNELL N, RABIE H, WHITELAW AC, DRAMOWSKI A. Neonatal and paediatric bloodstream infections: Pathogens, antimicrobial resistance patterns and prescribing practice at Khayelitsha District Hospital, Cape Town, South Africa. *SAMJ South African Medical Journal* 2018; **108**(2):99-104.
- 18. DENTI P, GARCIA-PRATS AJ, DRAPER HR, WIESNER L, WINCKLER JL, THEE S, DOOLEY KE, SAVIC R M, MCILLERON H, SCHAAF HS, HESSELING AC. Levofloxacin Population Pharmacokinetics in South African Children treated for multidrug-resistant tuberculosis. *Antimicrobial Agents and Chemotherapy* 2018; **62**(2):e01521-17.
- 19. DHANSAY MN. Conventional Indicators of the Burden of Young Child Malnutrition: time for a rethink?. *South African Journal of Clinical Nutrition* 2018; **31**(2):4-5.
- 20. DODD PJ, YUEN CM, BECERRA MC, REVILL P, JENKINS HE, SEDDON JA. Potential effect of household contact management on childhood tuberculosis: a mathematical modelling study. *Lancet Global Health* 2018; **6**:e1329-38.
- 21. DONALD PR, DIACON AH, LANGE C, DEMERS A, VON GROOTE-BIDLINGMAIER F, NARDELL EA. Droplets, dust and guinea pigs: an historical review of tuberculosis transmission research, 1878-1940. *International Journal of Tuberculosis and Lung Disease* 2018; **22**(9):972-982.
- 22. DOWLING WB, VELDSMAN KA, KATUSIIME MGK, MARITZ J, BOCK PA, MEEHAN S, VAN SCHALKWYK M, COTTON MF, PREISER W, VAN ZYL GU. HIV-1 RNA testing of pooled dried blood spots is feasible to diagnose acute HIV infection in resource limited settings. Southern African Journal of Infectious Diseases (Formerly Southern African Journal of Epidemiology and Infection) 2018; 33(2):50-53.
- 23. DRAMOWSKI A, LLOYD LG, BEKKER A, HOLGATE SL, AUCAMP M, REDDY K, FINLAYSON H. Neonatal listeriosis during a countrywide epidemic in South Africa: A tertiary hospital's experience. *SAMJ South African Medical Journal* 2018; **108**(10):818-827.
- 24. DU PREEZ K, DU PLESSIS L, O'CONNELL N, HESSELING AC. Burden, spectrum and outcomes of children with tuberculosis diagnosed at a district-level hospital in South Africa. . *International Journal of Tuberculosis and Lung Disease* 2018; **22**(9):1037-1043.
- 25. DU PREEZ K, SCHAAF HS, DUNBAR R, WALTERS E, SWARTZ A, SOLOMONS RS, HESSELING AC. Complementary surveillance strategies are needed to better characterise the epidemiology, care pathways and treatment outcomes of tuberculosis in children. *BMC Public Health* 2018; 18:397.
- 26. DUNBAR R, NAIDOO P, BEYERS N, LANGLEY I. Improving rifampicin-resistant tuberculosis diagnosis using Xpert® MTB/RIF: modelling interventions and costs. *International Journal of Tuberculosis and Lung Disease* 2018; **22**(8):890-898.
- 27. DZIKITI M, ESSER MM, SLOGROVE AL, BETTINGER JA, MACHEKANO R, COTTON MF. Early Breastfeeding Cessation Among HIV-Infected and HIV-Uninfected Women in Western Cape Province, South Africa. *AIDS and Behavior* 2018; **2018**(22):114-120.

- 28. ENGELBRECHT JG, MUKINDA FK, GREEN BM, SKINNER D. How clinicians experience a simulated antiretroviral therapy adherence exercise: A qualitative study. *African Journal of Primary Health Care and Family Medicine* 2018; **10**(1):a1836.
- 29. FLOYD S, AYLES H, SCHAAP AB, SHANAUBE K, MACLEOD D, PHIRI M, GRIFFITH S, BOCK PA, BEYERS N, FIDLER S, HAYES R. Towards 90-90: Findings after two years of the HPTN 071 (PopART) cluster-randomized trial of a universal testing-and-treatment intervention in Zambia. *PLoS One* 2018; **13**(8):e0197904.
- 30. FREGONESE F, AHUJA SD, AKKERMAN OW, ARAKAKI-SANCHEZ D, AYAKAKA I, BAGHAEI P, BANG D, BASTOS M, BENEDETTI A, BONNET M, CATTAMANCHI A, CEGIELSKI P, CHIEN J-Y, GARCIA-PRATS AJ, SCHAAF HS, ET AL. Comparison of different treatments for isoniazid-resistant tuberculosis: an individual patient data meta-analysis. *Lancet Respiratory Medicine* 2018; **6**:265-275.
- 31. FRIGATI LI, ARCHARY M, RABIE H, PENAZZATO M, FORD N. Priorities for Decreasing Morbidity and Mortality in Children with Advanced HIV disease. *Clinical Infectious Diseases* 2018; **66**(Suppl 2):S147-S151.
- 32. FRIGATI LJ, BEKKER A, STROBELE S, GOUSSARD P, SCHAAF HS. Culture-confirmed tuberculosis in South African infants younger than 3 months of age: clinical presentation and management of respiratory complications. *Pediatric Infectious Disease Journal* 2018; **1**(1):1.
- 33. FRY S, BARNABAS SL, COTTON MF. Update on trends in childhood tuberculosis. *CURRENT OPINION IN PEDIATRICS* 2018; **30**(1):152-160.
- 34. GARCIA-BASTEIRO AL, SCHAAF HS, DIEL R, MIGLIORI GB. Adolescents and young adults: a neglected population group for tuberculosis surveillance. *European Respiratory Journal* 2018; **51**:1800176.
- 35. GARCIA-PRATS AJ, DRAPER HR, FINLAYSON H, WINCKLER JL, BURGER A, FOURIE B, THEE S, HESSELING AC, SCHAAF HS. Clinical and cardiac safety of long-term levofloxacin in children treated for multidrug-resistant tuberculosis. *Clinical Infectious Diseases* 2018; **67**(11):1777-1780.
- 36. GARCIA-PRATS AJ, ROSE PC, DRAPER HR, SEDDON JA, NORMAN J, MCILLERON HM, HESSELING AC, SCHAAF HS. Effect of Co-Administration of Lidocaine on the pain and pharmacokinetics of intramuscular amikacin in children with multidrug-resistant tuberculosis: A randomized crossover trial. *Pediatric Infectious Disease Journal* 2018; **37**(12):1199-1203.
- 37. GARCIA-PRATS AJ, SVENSSON EM, WELD E, SCHAAF HS, HESSELING AC. Current status of pharmacokinetic and safety studies of multidrug-resistant tuberculosis treatment in children. *International Journal of Tuberculosis and Lung Disease* 2018; **22**(5):s15-s23.
- 38. GLANZMANN B, ABRAHAM DR, MOLLER M, GLASHOFF RH, VAN COLLER A, UREN CP, DURRHEIM GA, URBAN MF, VAN HELDEN EG, ESSER MM, RICE GI, CROW YJ, KINNEAR CJ. Variable expression in SAMHD1-Associated familial aicardi-Goutieres syndromes. *Current Allergy and Clinical Immunology* 2018; **31**(4):194-198.
- 39. GOUSSARD P. The role of bronchoscopy in wheezing preschool children. *Current Allergy and Clinical Immunology* 2018; **31**(2):75-79.
- 40. GREEN L, GOUSSARD P, VAN ZYL A, KIDD M, KRUGER M. Predictive Indicators to Identify High-Risk Paediatric Febrile Neutropenia in Paediatric Oncology Patients in a Middle-Income Country. *JOURNAL OF TROPICAL PEDIATRICS* 2017; **64**(1):395-402.
- 41. GROBBELAAR M, VAN TOORN R, SOLOMONS RS. Lumbar cerebrospinal fluid evolution in childhood tuberculous meningitis. *JOURNAL OF CHILD NEUROLOGY* 2018; **33**(11):700-707.
- 42. HARAUSZ EP, GARCIA-PRATS AJ, LAW S, SCHAAF HS, KREDO T, SEDDON JA, MENZIES D, TURKOVA A, ACHAR J, AMANULLAH F, BARRY PM, BECERRA C, CHAN ED, ET AL, HESSELING AC. Treatment and outcomes in children with multidrug-resistant tuberculosis: A systematic review and individual patient data meta-analysis. *PLoS Medicine* 2018; **15**(7):e1002591.

- 43. HARGREAVES JR, KRISHNARATNE S, MATHEMA HT, LILLESTON PS, SIEVWRIGHT K, MANDLA NF, MAINGA T, VERMAAK R, PIWOWAR-MANNING E, SCHAAP AB, DONNELL D, AYLES H, HAYES R, HODDINOTT G, BOND V, STANGL A. Individual and community-level risk factors for HIV stigma in 21 Zambian and South African communities: analysis of data from the HPTN071 (PopART) study. *AIDS* 2018; **32**(6):783-793.
- 44. HESSELING PB, AFUNCHWI G, NJAMSHI V, KIMBI C, NFOR P, NJODZEKA B, WHARIN P, KOUYA F, DRAPER HR. Burkitt lymphoma (BL): effect of cyclophosphamide (CPM) on the menarche and fertility rate in 113 Cameroonian patients. *BRITISH JOURNAL OF HAEMATOLOGY* 2018; **182**(Suppl 1):7.
- 45. HESSELING PB, KOUYA F, KATAYI E, MBAH G, WHARIN P. Burkitt's lymphoma: The prevalence of HIV/AIDS and the outcome of treatment. *SAMJ South African Medical Journal* 2018; **108**(2):84-85.
- 46. HIRSCH-MOVERMAN Y, MANTELL JE, LEBELO L, WYNN C, HESSELING AC, HOWARD AA, NACHMAN S, FREDERIX K, MAAMA LB, EL-SADR W. Tuberculosis preventive treatment preferences among care givers of children in Lesotho: a pilot study . *International Journal of Tuberculosis and Lung Disease* 2018; 22(8):858-862.
- 47. HODDINOTT G, HESSELING AC. Social science is needed to understand the impact of paediatric MDR-TB treatment on children and their families. *International Journal of Tuberculosis and Lung Disease* 2018; **22**(1):4.
- 48. HODDINOTT G, MYBURGH H, NDUBANI R, MANTANTANA J, THOMAS A, MBEWE M, AYLES H, BOCK PA, SEELEY J, SHANAUBE K, HARGREAVES JR, BOND V, REYNOLDS LJ. Households, fluidity, and HIV service delivery in Zambia and South Africa an exploratory analysis of longitudinal qualitative data from the HPTN 071 (PopART) trial. *Journal of the International AIDS Society* 2018; **21**(s4):e25135.
- 49. HODDINOTT G, STAPLES S, BROWN RA, SIMWINGA M, MUBEKAPI-MUSADAIDZWA C, HESSELING AC, HENDRICKS GA, DE KOKER P, MCKENNA L. Community engagement for paediatric MDR-TB clinical trials: principles to support ethical trial implementation. . *International Journal of Tuberculosis and Lung Disease* 2018; **22**(5):s40-s45.
- 50. HONETH IE, LAUGHTON B, SPRINGER PE, COTTON MF, PRETORIUS C. Diagnostic accuracy of the Molteno Adapted Scale for developmental delay in South African toddlers. *Paediatrics and International Child Health* 2018; **1**(1):1-7.
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# Journal Articles (NON-subsidised)

- 130. DE WILDE C, DU PLESSIS A, GOUSSARD P. Unusual presentation of pulmonary hydatidosis mimicking thoracic malignancy in a paediatric South African patient. *Journal of the Belgian Society of Radiology* 2018; **102**(1):70, 1-4.
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# **Chapters in Books**

- 133. KRUGER M. Childhood Cancers and its warning signs. In: de Villiers M (ed.) *Handbook of Oncology*, MIMS, Saxonworld, South Africa, 2018: 119-122.
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- 135. SCHAAF HS, MARAIS BJ, CARVALHO I, SEDDON JA. Challenges in childhood tuberculosis. In: Migliori GB (ed.) *Tuberculosis*, European Respiratory Society, Sheffield, United Kingdom, 2018: 234-262.

# **CENTRES**

# THE FAMILY CLINICAL RESEARCH UNIT (FAMCRU)

**Director: Prof MF Cotton** 



In 2018, FAMCRU conducted 20 clinical trials and research studies, two thirds of which were in paediatric participants. These studies are conducted within the NIH funded IMPAACT (International Maternal Pediatric Adolescent AIDS Clinical Trials Group) and ACTG (AIDS Clinical Trials Group) networks, the European PENTA-ID (Paediatric European Network for Treatment of AIDS – Infectious Diseases) network, pharmaceutical funded trials and investigator driven trials. We screened 564 potential participants and enrolled 322. There were 6041 clinic visits comprising 4179 scheduled visits and 1862 unscheduled during 2018. Additions to the staff complement include 2 pharmacists, 2 doctors, one 1 administrative and a human resources assistant.

# 1. RESOURCE SHARING AND COLLABORATION

Dr Barbara Laughton conducts the following neurodevelopment studies:

1.1 The Association of Perinatal HIV infection and Hearing Loss in Children of Cape Town, South Africa.

Funding: R01DC015984, NIH

Collaborators: P Torre (PI) San Diego State University, Tj Yao, Harvard University, Andre van Der Kouwe Massachussets General Hospital,

Ernesta Meintjes UCT, Marcin Jankiewicz UCT.

1.2 Neuroimaging and gut microbiome markers of development in HIV-exposed uninfected infants.

Funding: R01HD093578, NIH

Collaborators: MJ Holmes (UCT), M Kaba (UCT), A van der Kouwe (MGH), F Little (UCT)

1.3 Neonatal Imaging as a marker of neurodevelopment and predictor of cognitive

Performance in infants exposed to HIV and ART in utero and perinatally.

Funding: R01 HD085813, NIH

Collaborators: A van der Kouwe (MGH), E Meintjes (UCT), S Jacobson

(Wayne State University), J Jacobson (Wayne State University), F Little (UCT)

1.4 Latent reservoir characterization and correlations with neurocognitive function.

Funding: R01MH105134, NIH

Collaborators: Virology, Immunology, Neuroimaging at UCT

# 2. ACHIEVEMENTS AND HIGHLIGHTS

# 2.1 Appointments

**1.1.1** Marije van Schalkwyk – voting member of WHISC (Women's Health Inter-Network Scientific Committee)

- **1.1.2** Shaun Barnabas member of the ACTG CURE Committee
- **1.1.3** Lynne Cornellisen member of the South African Board of Trustees and Chair of the Global Programs Committee for Mothers 2 Mothers
- **1.1.4** Sonja Pieterse Pharmacy Manager at FAMCRU
- **1.1.5** Mark Cotton serves on the IMPAACT CURE Committee
- **1.1.6** Joan Coetzee serves on:
  - Protocol Development and Implementation Sub-committee (PDISC) Cochair
  - Scientific Agenda Steering Committee (SASC) SMCCC representative Site Management Clinical Care Committee (SMCCC) member Field Representative Working Group (FRWG) member
- **2.1.7** Marie Theunissen serves as a member of the International Community Advisory Board
- **2.1.78** Steve Innes serves on the IMPAACT Complications Scientific Committee

# 2.2 Grants / Awards

- **1.2.1** Gerhard Walzl won GOLD medal MRC reward for Research Excellence
- **1.2.2** 4 CAB members were awarded the Bridgitte Murtagh award for continuous dedication through-out the year as a CAB member
- 1.2.3 Shalena Naidoo received the International AIDS Society and ANRS Dominique Dormont award for young researchers working on chronic conditions interfacing between HIV. The prize highlights researchers who demonstrate originality, rationale, quality and a multidisciplinary and integrative approach in HIV research for her poster "Persistence of myeloid cell-associated inflammation in HIV-infected children after 8 years on early initiated therapy the key role players in HIV persistence?"

#### 2.3 Educational achievements

- **1.3.1** Dr Lindie Rossouw: BTh Honours, Dec 2018
- **1.3.2** Dr Samantha Fry MMed Paediatrics

### 3. TRAINING AND TEACHING

# 3.1 Academic meetings

- 3.1.1 Quarterly SUN CTU meetings
- 3.1.2 Weekly SUN CTU academic presentations
- 3.1.3 Participation in the weekly Tygerberg Paediatric Infectious Diseases Meeting
- 3.1.4 Regular representation by Samantha Fry and Marije van Schalkwyk at the 2-monthly Tygerberg subdistrict HIV and TB meeting for community clinics and the quarterly Metro East HIV and TB meeting.

#### 3.2 Courses Attended

- **1.2.1** Neuropsychs Tester-Griffiths III Training Mrs B Fanqa (certified course required to test on the Griffiths)
- **1.2.2** Alarm Distress Baby Scales Scoring Dr B Laughton (certified course require to test on the Griffiths)
- 1.2.3 National Radiology Refresher Course, August M van Schalkwyk
- **1.2.4** BLS and ACLS course, Dec M van Schalkwyk, L Cornellison
- 1.2.5 APLS, Advanced Life Support Group, 15 March 2018 Magdel Rossouw
- **1.2.6** Performance Management Training, Stellenbosch University, March 2018 Jeanne Louw

- **1.2.7** SU Staff HIV Counselling and Testing Short course, 25 27 Sep 2018: Veronica
  - Matroos & Shiella Dyan
- **1.2.8** Magdel Rossouw: APLS, Advanced Life Support Group, 15 March 2018
- **1.2.9** Jeanne Louw, Performance Management Training, Stellenbosch University, March 2018
- 1.2.10 Breastfeeding Seminar, Western Cape Government (Health) 3 Aug 2018 Dr Magdel Rossouw, Dr Lindie Rossouw, Phoebe Ganjana, Nokuza Bonani, Leonie Groepies, Sylvia Pieterse, Desmien Bagus, Bongiwe Makeleni, Veronica Matroos
- 1.2.11 Stellenbosch University Annual Academic Day 29 Aug 2018:
  Prof Gerhard Theron, Jeanne Louw, Dr Magdel Rossouw, Dr Lindie Rossouw,
  Phoebe Ganjana, Nokuza Bonani, Wendy Adendorff, Leonie Groepies, Sylvia
  Pieterse, Desmien Bagus, Bongiwe Makeleni, Veronica Matroos, Shiella Dyan
- 1.2.12 Paediatrics and Child Health Academic Day of Excellence, Stellenbosch University 7 Nov 2018:
  Dr Magdel Rossouw, Dr Lindie Rossouw, Phoebe Ganjana, Nokuza Bonani, Wendy Adendorff, Leonie Groepies, Sylvia Pieterse, Desmien Bagus, Bongiwe Makeleni, Veronica Matroos, Shiella Dyan
- **1.2.13** SU Staff HIV Counselling and Testing Short course, 25 27 Sep 2018: Veronica Matroos & Shiella Dyan

# Staff training

- 1.2.14 GCP Anthony Hitchcock, Sharon Kruger
- 1.2.15 Neurodevelopmental:
  - 1.2.15.1 Preparation and positioning of newborns for un-sedated MRI scans for research study F Lindani, T Hamana, S Fry
  - 1.2.15.2 Fagan test of infant intelligence (trained to test) Mrs B Fanqa
  - 1.2.15.3 WHO Infant Dietary Survey HBS study team
  - 1.2.15.4 Dubowitz Neurological Assessment of the Newborn Dr S Fry
  - 1.2.15.5 HIV adolescent workshop, 10-12 Oct P Zuidewind
  - 1.2.15.6 Performance Management training, 9 March A J van Rensburg
  - 1.2.15.7 Adherence counselling training, Sept T Ncgaba
  - 1.2.15.8 HIV Counselling, 25 27 Sep Laurette Siphambo (Counsellor)
  - 1.2.15.9 Nuts and Bolts and Safety training, 3 5 Dec Doreen Willemse, Dr Olivia Arendse, Dr Amy Slogrove
  - 1.2.15.10 IATA Training, 15 -17 Aug Anthony Hitchcock

# 1.2.16 IMPAACT / ACTG:

- 1.2.16.1 GCP lectures Developing SOPs and Recruitment and Retention of Research Participants.
- 1.2.16.2 Mentoring 2 Study Coordinators
- 1.2.16.3 Staff training in the consent process
- 1.2.16.4 CAB leadership training for the CAB executive committee

# 1.3 Undergraduate Teaching

- **1.3.1** Samantha Fry assisted with 3<sup>rd</sup> year undergraduate paediatric clinical teaching
- **1.3.2** Shaun Barnabas assisted with 3<sup>rd</sup> year undergraduate paediatric lectures
- 1.3.3 Shaun and Samantha assist with middle rotation and SI clinical examinations

#### 1.4 International Students

# **1.4.1** FAMCRU. together with HOPE hosted 4 international students completing their clinical elective rotation

# 4. CONFERENCES: PRESENTATIONS AND ATTENDANCE

| Name       | Conference                                                             | Date<br>(2018) | Attendee / oral presentation / poster presentation / key note speaker / guest speaker / plenary / session chair / other |
|------------|------------------------------------------------------------------------|----------------|-------------------------------------------------------------------------------------------------------------------------|
| B Laughton | 41 <sup>st</sup> Paediatrics refresher course UCT                      | 15 Feb         | Attendee                                                                                                                |
| B Laughton | 10 <sup>th</sup> International workshop on HIV Pediatrics, Amsterdam   | July           | Attendee                                                                                                                |
| S Naidoo*  | 10 <sup>th</sup> International workshop on HIV<br>Pediatrics Amsterdam | July           | Oral presentation of work from study at FAMCRU                                                                          |
| I Familiar | 10 <sup>th</sup> International workshop on HIV<br>Pediatrics Amsterdam | July           | Oral presentation of work from Multisite study at FAMCRU                                                                |
| E Nwosu    | 10 <sup>th</sup> International workshop on HIV<br>Pediatrics Amsterdam | July           | Poster presentation of work from study at FAMCRU                                                                        |
| F Mberi    | 10 <sup>th</sup> International workshop on HIV<br>Pediatrics Amsterdam | July           | Poster presentation                                                                                                     |
| B Laughton | HEU workshop                                                           | July           | Attendee                                                                                                                |
| B Laughton | Adolescent workshop                                                    | July           | Attendee                                                                                                                |
| B Laughton | 22 <sup>nd</sup> International AIDS conference                         | July           | Attendee                                                                                                                |
| A Ibrahim  | 22 <sup>nd</sup> International AIDS conference                         | July           | Poster presentation of work from study at FAMCRU                                                                        |
| FMberi     | 22 <sup>nd</sup> International AIDS conference                         | July           | Poster presentation of work from study at FAMCRU                                                                        |
| JS Madzime | 22 <sup>nd</sup> International AIDS conference                         | July           | Poster presentation of work from study at FAMCRU                                                                        |
| EC Nwosu   | 22 <sup>nd</sup> International AIDS conference                         | July           | Poster presentation of work from study at FAMCRU                                                                        |
| B Laughton | CIPHER Stakeholder Meeting                                             | July           | Attendee                                                                                                                |
| T Hamana   | 22 <sup>nd</sup> International AIDS conference                         | July           | Poster presentation of work from study at FAMCRU                                                                        |
| M. Cotton  | 22 <sup>nd</sup> International AIDS conference                         | July           | Oral presentation: Pharmacokinetics, safety, & efficacy of                                                              |

|                  |                                              |            | Internal Control of the Control of |
|------------------|----------------------------------------------|------------|------------------------------------|
|                  |                                              |            | bictegravir, emtricitabine         |
|                  |                                              |            | & tenofovir alafenamide            |
|                  |                                              |            | (B/F/TAF) single-tablet            |
|                  |                                              |            | regimen in HIV-1-                  |
|                  |                                              |            | infected children (6 to <          |
|                  |                                              |            | 12 years)                          |
| Keri Woods       | 62 <sup>nd</sup> AAD Stellenbosch University | August     | Poster presentation of             |
|                  |                                              |            | work from study at                 |
|                  |                                              |            | FAMCRU                             |
| Shalena Naidoo   | 62 <sup>nd</sup> AAD Stellenbosch University | August     | Oral presentation of work          |
|                  |                                              |            | from study at FAMCRU               |
| B Laughton       | SAPA conference                              | September  | Poster presentation                |
|                  |                                              |            | X 2                                |
| Amy Graham       | World Audiology Congress                     | October    | Poster presentation of             |
|                  |                                              |            | work from study at                 |
|                  |                                              |            | FAMCRU                             |
| P Torre          | World Audiology Congress                     | October    | Oral presentation of work          |
|                  |                                              |            | from study at FAMCRU               |
| H Elliot         | World Audiology Congress                     | October    | Poster presentation of             |
|                  |                                              |            | work from study at                 |
|                  |                                              |            | FAMCRU                             |
| A Janse van      | IMPAACT 2018                                 | June       | Attendee                           |
| Rensburg         | PENTA 20 (Odyssey)                           | 17-22 July | Attended                           |
|                  | HIV Paediatric workshop                      | 20-21July  |                                    |
| M Cotton, S      | Annual AIDS Clinical Trials Group            | June       | Attendee                           |
| Barnabas, G      | (ACTG) Scientific Retreat Meeting            |            |                                    |
| Fourie           |                                              |            |                                    |
| Joan Coetzee     | Annual ACTG Network Meeting                  | June       | Attendee/ Facilitator of           |
|                  |                                              |            | PDISC committee fair               |
| M Cotton         | CROI 2018, Boston Massachusetts              | March      |                                    |
| J Louw;          | IMPAACT Annual Meeting                       | June       | Attendees                          |
| G Theron, M      |                                              |            |                                    |
| Cotton, S Fry, S |                                              |            |                                    |
| Barnabas, G      |                                              |            |                                    |
| Fourie, A        |                                              |            |                                    |
| Fourie, M        |                                              |            |                                    |
| Smuts, J         |                                              |            |                                    |
| Coetzee          |                                              |            |                                    |

# 5. PUBLICATIONS

- 5.1 Laughton B, Cornell M, Kidd M, Springer PE, Dobbels EFM, Rensburg AJV, Otwombe K, Babiker A, Gibb DM, Violari A, Kruger M, Cotton MF. Five year neurodevelopment outcomes of perinatally HIV-infected children on early limited or deferred continuous antiretroviral therapy. Journal of the International AIDS Society. 2018; 21(5):e25106. PubMed [journal] PMID: 29722482, PMCID: PMC5932637
- **5.2** Thomas A, Dobbels EFM, Springer PE, Ackermann C, Cotton MF, Laughton B. Favourable outcome in a child with symptomatic diagnosis of Glutaric aciduria type 1

- despite vertical HIV infection and minor head trauma. Metabolic brain disease. 2018; 33(2):537-544. PubMed [journal] PMID: 29427049
- 5.3 Robertson FC, Holmes MJ, Cotton MF, Dobbels E, Little F, Laughton B, van der Kouwe AJW, Meintjes EM. Perinatal HIV Infection or Exposure is associated with low N-Acetylaspartate and Glutamate in Basal Ganglia at age 9 but not 7 years. Front. Hum. Neurosci. 12:145. Doi 10.3389/fnhum.2018.00145
- 5.4 Chernoff MC1, Laughton B2, Ratswana M3, Familiar I4, Fairlie L3, Vhembo T5, Kamthunzi P6, Kabugho E7, Joyce C8, Zimmer B9, Ariansen JL10, Jean-Philippe P11, Boivin MJ. Validity of Neuropsychological Testing in Young African Children Affected by HIV.J Pediatr .Infect Dis. 2018 Sep;13(3):185-201. doi: 10.1055/s-0038-1637020.
- **5.5** Fry S, Barnabas S, and Cotton MF, Update on trends in childhood tuberculosis. Curr Opin Pediatr 30 (2018) 152-160.
- 5.6 Innes S, van der Laan L, Anderson PL, Cotton M, and Denti P, Can We Improve Stavudine's Safety Profile in Children? Pharmacokinetics of Intracellular Stavudine-Triphosphate with Reduced-Dosing. Antimicrob Agents Chemother (2018).
- **5.7** Slogrove AL, Powis KM, Bettinger JA, and Cotton MF, Considerations in evaluating infectious morbidity and mortality in HIV-exposed uninfected infants. AIDS 32 (2018) 2855-2856.
- **5.8** Veldsman KA, Maritz J, Isaacs A, Katusiime MG, Janse Van Rensburg A, Laughton B, Mellors JW, Cotton MF, and Van Zyl GU, Rapid decline of HIV-1 DNA and RNA in infants starting very early ART may pose a diagnostic challenge. AIDS (2018).
- 5.9 Zunza, M. Esser, A. Slogrove, J.A. Bettinger, R. Machekano, M.F. Cotton, and C. Mother-Infant Health Study Project Steering, Early Breastfeeding Cessation Among HIV-Infected and HIV-Uninfected Women in Western Cape Province, South Africa. AIDS Behav (2018).

# 6. COMMUNITY ENGAGEMENT

#### **6.1 Community Advisory Board**

- **6.1.1** Combined Community Advisory Board conference 18 19 October 2018
- **6.1.2** Gala event for Community Advisory Board members
- **6.1.3** Establishment of the Katlegho Mosimangape award for young people belonging to the CAB and have proved to be dedicated in their role as community supporter and peer educator.
- **6.1.4** Mandela Day Contributed to TygerBear knit a teddy project

# 7. INTERNATIONAL VISITORS

# 5.1 Neurodevelopmental:

- 5.1.1 Dr Peter Torre from San Diego State University, PI on Audiology study
- 5.1.2 Dr Andre van der Kouwe from Massachusetts General Hosp (Harvard), Co-PI on Healthy Baby study
- **5.2** Cheryl Pikora, Medical monitor from Gilead, 16 Nov 2018
- **5.3** Philip Goulder Oxford University

# **DESMOND TUTU TB CENTRE (DTTC)**

**Director: Prof AC Hesseling** 

# **DESMOND TUTU TB CENTRE ANNUAL REPORT: 2018**



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#### LETTER FROM THE DIRECTOR

The past year was exciting and challenging for our centre. Large trials like HPTN 071 (PopArt) have come to an end, resulting in much smaller staff component and reduced funding.

At the same time, we have become more efficient, more internally collaborative and our three "research pillars" in our centre have become more integrated. We have pursued new endeavours, notably in paediatric therapeutics and in adult HIV prevention trials.

We continue to build capacity at all levels and have had a record 6 PhD students graduate from our centre during 2018. There is now a strong cadre of young and mid-level career researchers who are not only supporting our scientific agenda, but are mentoring and supporting other researchers.

We have established new local, South African, Africana and global collaborations and continue to be the global leader in paediatric tuberculosis clinical research.

We have opened two new research sites to support TB and HIV prevention trials, and have grown in key core support areas including regulatory, data, laboratory, logistics and communications.

We are thankful for a highly productive year and look forward to going from strength to strength in 2019.

Professor Anneke C. Hesseling

alus

Director: Desmond Tutu TB Centre

Distinguished Professor in Paediatrics and Child Health

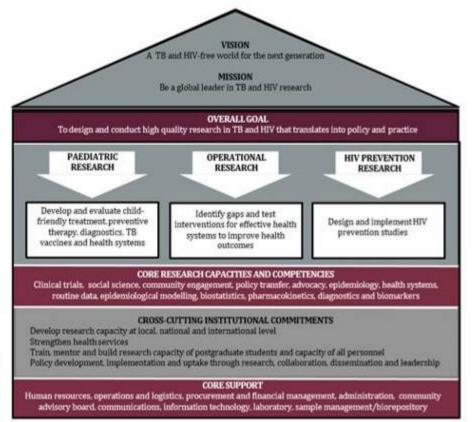
SARcHi Chair in Paediatric Tuberculosis

#### **DTTC STRATEGY HOUSE**



# Desmond Tutu TB Centre Strategy House







# **GLOSSARY OF TERMINOLOGY**

| ACTG       | AIDS Clinical Trials Group                                             |  |
|------------|------------------------------------------------------------------------|--|
| AE/AER/EAE | Adverse Event / Adverse Event Report / Expedited Adverse Event         |  |
| AIDS       | Acquired Immunodeficiency Syndrome                                     |  |
| ART/ARV    | Antiretroviral Therapy / Antiretroviral                                |  |
| ВМС        | BioMed Central                                                         |  |
| BMRC       | British Medical Research Council                                       |  |
| CAB/CAG    | Community Advisory Board/Community Advisory Group                      |  |
| CD4        | Cluster of Differentiation 4                                           |  |
| CDC        | US Centers for Disease Control and Prevention                          |  |
| CHIP       | Community HIV Care Providers                                           |  |
| CRS        | Clinical Research Site                                                 |  |
| CTU        | Clinical Trials Unit                                                   |  |
| CXR        | Chest X-ray                                                            |  |
| DAIDS      | Division of AIDS, NIAID                                                |  |
| DFID       | Department for International Development                               |  |
| DR-TB      | Drug-Resistant Tuberculosis                                            |  |
| DS-TB      | Drug-Susceptible Tuberculosis/Drug-Sensitive Tuberculosis              |  |
| DST        | Drug Susceptibility Testing                                            |  |
| EC         | Ethics Committee                                                       |  |
| DTTC       | Desmond Tutu TB Centre                                                 |  |
| EDCTP      | The European & Developing Countries Clinical Trials Partnership        |  |
| FAMCRU     | Family Clinical Research Unit                                          |  |
| FDA        | Food and Drug Administration                                           |  |
| FHI 360    | Family Health International and Academy for Educational Development    |  |
| FMHS       | Faculty of Medicines and Health Science                                |  |
| HIV        | Human Immunodeficiency Virus                                           |  |
| HPTN       | HIV Prevention Trials Network                                          |  |
| HR         | Human Resources                                                        |  |
| HREC       | Health Research Ethics Committee                                       |  |
| HTS        | HIV Testing Services                                                   |  |
| IMPAACT    | International Maternal Pediatric Adolescent AIDS Clinical Trials Group |  |
| IT         | Information Technology                                                 |  |
| LAB        | Laboratory                                                             |  |
| MA         | Master of Arts                                                         |  |
| MCC        | Medicines Control Council                                              |  |
| MDR-TB     | Multidrug-Resistant Tuberculosis                                       |  |
| MPH        | Master of Public Health                                                |  |
| MRC        | Medical Research Council                                               |  |
| MS         | Mass Spectrometry                                                      |  |
| MSF        | Médecins sans Frontières                                               |  |
| L          |                                                                        |  |

| M.tb   | Mycobacterium tuberculosis                                    |
|--------|---------------------------------------------------------------|
| NIAID  | National Institute of Allergy and Infectious Diseases         |
| NICHD  | National Institute of Child Health and Human Development      |
| NIH    | National Institutes of Health                                 |
| OR     | Operational Research                                          |
| ORAP   | Operational Research Assistance Project                       |
| PC     | Population Cohort                                             |
| PEPFAR | President's Emergency Plan for AIDS Relief                    |
| PHC    | Primary Health Care                                           |
| PI     | Principal Investigator                                        |
| PK     | Pharmacokinetics                                              |
| PMTCT  | Prevention of Mother To Child Transmission                    |
| RIF    | Rifampicin                                                    |
| RGMO   | Research grants management office                             |
| QA     | Quality Assurance                                             |
| SACEMA | South African Centre for Epidemiological Modelling & Analysis |
| STI    | Sexually transmitted infections                               |
| SU     | Stellenbosch University                                       |
| TBTC   | TB Trials Consortium                                          |
| VMMC   | Voluntary Male Medical Circumcision                           |
| ZAR    | South African Rand (currency)                                 |

#### **ORGANOGRAM**

#### SCIENTIFIC FOCUS AREAS

#### DTTC DIRECTOR: PROF AC HESSELING

#### Paediatric Research

#### Investigators

Prof Anneke Hesseling Prof Simon Schaaf

Prof Peter Donald

Prof Robert Gie

Dr Anthony Garcia-Pratts

Dr Jana Winckler

Dr Louvina van der Laan

Dr Elisabetta Walters

Dr Anne-Marie Demers

Dr Marieke van der Zalm

Dr Megan Palmer

Dr Sue Purchase

Prof Adrie Bekker

Dr Celeste de Vaal

Dr Jennifer Hughes

Dr Rolanda Croucamp

Dr James Seddon

Dr Elisa Lopez

#### Study Coordinators

Petra de Koker Jessica Workman Sharon Mbaba Margaret van Niekerk Adelaide Carelse Elise Batist Nicola James

Nurses

Counsellors / DOT workers

Staff component including support staff – n=84

# Operational Research

#### Investigators

Dr Karen du Preez

Dr Muhammad Osman

Dr Sue - Ann Meehan

Graduate Research Assistants

Research Apprentices

Staff component including support staff - n=7

#### HIV Prevention Research

#### Investigators

Prof Nulda Beyers Dr. Peter Bock

Project Managers/Clinicians

Dr Kerry Joubert Nomtha Mandla

Study Coordinators/Junior Project/District Managers

Vikesh Naidoo Dr. Gerald Maarman Yvonne Saunders Nozizwe Makola

#### Nurses

District Logistics Officers Counsellors Community Engagement Officer Community Mobilizers

Staff component including support staff -n= 36

#### Socio-behavoural Research

Investigator

Graeme Hoddinott

#### Research Coordinators

Lario Viljoen Constance Mubekapi-Musadaidzwa Hanlie Myburgh

#### Project Coordinators

Angelique Thomas Laing de Villiers Dillon Wademan Rozanne Casper

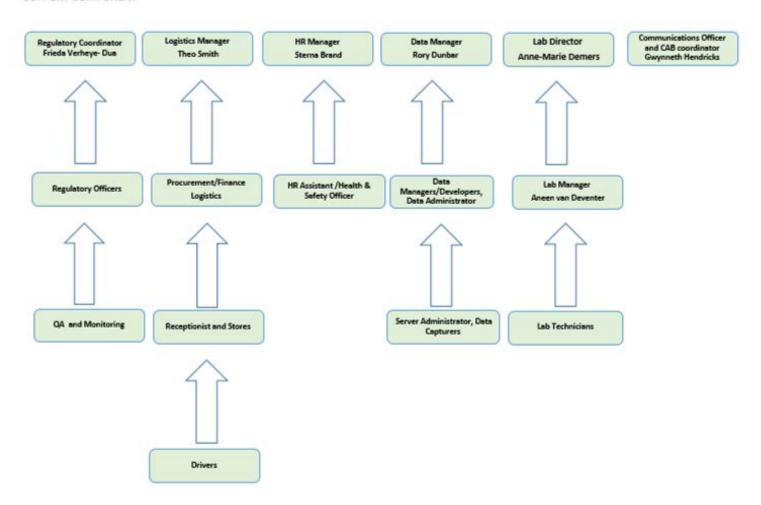
Graduate Research Assistants

Research Interns

Data Quality Assistants

Staff component including support staff - n=25

#### SUPPORT COMPONENT



DTTC TOTAL STAFF COMPONENT 152

#### **RESEARCH UPDATES**

# Research Focus Area One: Paediatric Tuberculosis

The DTTC initially has established its reputation based on its strong track record of clinical research in paediatric tuberculosis (TB). The DTTC is now the global leader in therapeutics for the prevention and treatment of TB in children, especially in the context of drug-resistant TB.

This paediatric TB research programme at the DTTC is led by Anneke Hesseling, centre director, SArCHI Chair in Paediatric Tuberculosis and Distinguished Professor in Paediatrics and Child Health, Stellenbosch University. Key focus areas include 1) therapeutics for Drug Susceptible (DS) and Drug Resistant TB (DR-TB) in children (leads; Anthony Garcia-Prats, Megan Palmer), 2) preventive trial for TB in children (lead: Anneke Hesseling, 3) Lung health, diagnostics and biomarkers in children (lead: Elisabetta Walters, Marieke van der Zalm and Anne-Marie Demers), 4) underpinned by in-depth epidemiological and operational research (leads: Karen du Preez, Muhammad Osman). Major areas of interest specifically include the evaluation of novel therapeutic strategies for Multidrug resistant TB (MDR-TB) in children, where it is a global leader in its field and is generating seminal data on an ongoing basis. The DTTC officially opened its upgraded state of the art paediatric pharmacokinetics clinical research unit at Brooklyn Chest Hospital (medical director: Tony Garcia-Prat) in July 2016 (refer to highlights, 2016), where it has been working since 2011. This unit has been expanded twice since then, reflecting the urgent need for clinical research in this domain, with 9 trials for antituberculosis treatment (including novel drugs like delamanid and bedaquiline) currently ongoing.

During 2018, the paediatric group (n=84 personnel) has specifically actively pursued linking and collaborating with the Sociobehavioural science group at DTTC (lead: Graeme Hoddinott), and on high quality operational research. The group has also expanded to collaborations at the Tygerberg campus on the role of viral pathogens in TB disease pathogenesis in children (Professor Gert van Zijl, Medical Virology), antimicrobial resistance in children and the biome and Medical Microbiology (Professor Andrew Whitelaw) and at the Stellenbosch campus, including health economics (Professor Ronelle Burger) mathematical modelling (SACEMA, Dr. Alex Welte, Eduard Grebe and Juliet Pulham), and basic scientists including biochemists (Professor Jacky Snoep), to support its expanding research agenda.

Expanding national collaboration have included Shandukani (Dr. Lee Fairlie, WHRI), and PHRU (Professor Neil Martinson) and new international collaboration have included the Uppsala pharmacometrics modelling group and Professor Tony Hu, Arizona State University and Professor Bob Husson (Boston Children's). The DTTC is also increasingly collaborating with other African research groups including in Uganda, Zambia, Mozambique and Namibia. During 2018, one of the pediatric team clinicians, Dr Karen du Preez, has been selected and trained as an epidemiological TB program review consultant for the World Health Organization, strengthening our existing collaboration to support pediatric TB at both national and international level.

The DTTC paediatric group is a well-performing clinic site for the DAIDS-funded International Maternal, Paediatric, Adolescent AIDS Clinical Trials (IMPAACT) where it is currently conducting 5 IMPAACT TB trials and also for the US Centres for Disease Control (CDC) TB Clinical Trials Consortium (TBTC), where it leads Study 35, a TB prevention trial of rifapentine and isoniazid, in children. The paediatric TB research group at the DTTC has been exceptionally productive during 2018. They have graduated 2 Master's degree and 2 PhD students, with 2 new PhD students registered, 1 postdoctoral fellow from Spain (Elise Lopez), and a BMRC-funded research fellow from Imperial College London, James Seddon (who has received an extraordinary appointment at the DTTC as Senior Researcher).

The group's research has resulted in >50 publications in international peer-reviewed journals (Appendix I), participation in several national and international TB treatment guideline meetings, including with WHO, and with the SA National DOH.

The group has had 3 major international grants awarded, and several local grants.

The group has embarked on new collaborations with Imperial Collage London, Rutgers University, BJMC, Pune, India and La Salle Medical Health Sciences Institute College of Medicine, De La Salle University Medical Center, the Philippines. The group has also developed new national partners in South Africa, and new collaborative partners in Africa (Uganda and Zambia).

The therapeutics group has been shaping the science and implementation of paediatric TB treatment trials through leadership in the National Institutes of Health (NIH) funded IMPAACT and the CDC TBTC trial consortia.

The DTTC paediatric group hosted its 12<sup>th</sup> International TB Training Course at Goudini very successfully, with 50 international participants. The DTTC paediatric group alone contributed 13 presentations to the 49th annual International Union against Tuberculosis and Long Disease conference in The Hague, with 7 oral presentations, including a later breaker session.

# Therapeutic trials: prevention of drug-susceptible TB

**IMPAACT P1078**: This IMPAACT-funded network multisite randomized controlled double blinded trial evaluates the safety and tolerability of isoniazid ante- vs. post-partum in HIV- infected pregnant women. DTTC was the last site to join this trial as a newly funded IMPAACT site, in 2015. Twelve women were enrolled and follow-up visits completed July 2017. The overall trial was closed to accrual in 2016 and follow-up (740 maternal-infant pairs) was completed in September 2017 and the data is being disseminated.

**TBTC Study 35**: This multisite study, funded by the CDC TBTC (PI: Anneke Hesseling,) will evaluate the optimal dosing and safety of the novel 12 dose combination regimen of rifapentine and isoniazid, shown to be efficacious in prevention of TB in adults and children. The protocol has been in development for 4 years and has been finalized and approved by the Medicines Control Council (MCC) and local ethics committees. A novel fixed dose score paediatric formulation has been developed for the trial. The database is in development and the trial is expected to open in 2019 under an FDA IND.

# Therapeutic trials: treatment of drug-susceptible TB

SHINE: (Site PI: Anneke Hesseling). The trial PI and sponsor is the Medical Research Council Clinical Trials Unit (MRC CTU) at University College London (PI: Diane Gibb), with the funders: BMRC, Wellcome Trust, DFID. This multi-site open-label international trial is the first ever to evaluate the efficacy and safety of 4 vs. the standard 6-month WHO-recommended regimen for treatment shortening of non-severe DS-TB in children. Recruitment was completed in July 2018, with 1204 children enrolled in South Africa, Zambia, Uganda and India. DTTC enrolled a total of 315 children. Nested Pharmacokinetics (PK) sampling was completed in 31 children. This trial, if successful, will have a major impact on the current long treatment regimens for paediatric TB, which is usually paucibacillary (smear negative). DTTC has led nested qualitative work to evaluate the acceptability and palatability of the new WHO-endorsed fixed dose combination formula used in the trial. DTTC is also coordinating the process for central expert review of the Chest X-rays (CXR) taken on the SHINE trial – an undertaking which will involve review of approximately 3000 CXR images and will form an important part of the final adjudication of certainty of TB diagnosis and classification of outcomes on the trial. The evaluation of baseline CXRs by central experts is due to be completed in 2019. **Project P1026s:** Pharmacokinetic properties of antiretroviral and related drugs during pregnancy and postpartum (Site PI: Anneke Hesseling). The Phase IV, observational Pharmacokinetic (PK) study started in November 2017. The primary outcome is to describe the pharmacokinetic parameters during pregnancy of selected ARV drugs currently used in the clinical care of 25 HIV-infected pregnant women, and to compare these parameters to a) historical pharmacokinetic data from nonpregnant women, and b) postpartum pharmacokinetic data from the same women in the study cohorts. In 2018, one of the study clinicians addressed the clinicians at Brooklyn Chest Hospital and gave an overview of the studies currently run at the unit. This oral presentation was held at the clinicians' weekly clinical meeting at the hospital. The study clinician took this opportunity to meet

the newly appointed CEO of the hospital, Waheeda Sonnie, who showed her interest in the unit's work. To date, 8 women have enrolled in the study at the DTTC in the second-line TB drug arm of the protocol. This will provide critically needed information on the pharmacokinetics in pregnant women of medications used for MDR-TB. A follow-up study to 1026s, which will include first and second-line TB drug arms, is in development.

**IMPAACT P1101:** (Site PI: Anneke Hesseling) this IMPAACT-funded multisite network phase I/II trial is evaluating the PK and safety of raltegravir, a new integrase inhibitor, in combination with first-line TB treatment, including rifampicin. Children must be HIV-infected, on TB treatment but not currently on ARVs. This patient group has been difficult to enroll in Cape Town, given good HIV prevention and treatment services in children. Three participants have been enrolled to date and interim data analysis is ongoing for the overall trial (5 South African sites). A new cohort of children <2 years of age has been opened for the trial.

OptiRif Kids: (PI Anneke Hesseling) this phase I/II open label multi-cohort dose escalation study to evaluate the pharmacokinetics and safety of increased doses of rifampicin in HIV-negative children with TB disease is funded by TB Alliance. The study utilizes a dose escalation approach in a maximum of 5 dosing cohorts of 20 children aged 0-12 years in each cohort. This trial includes collaboration with the University of Cape Town Clinical Pharmacology Division, Radboud University, Nijmegen, and Uppsala University, Sweden as well as with the Shandukani Research Site, Wits Health Research Institute. This study aims to establish the dose of RMP in children required to match RMP exposure targets in adults, using a 35-40 mg/kg dose, over 15 days. In 2018, Cohorts 1 and 2 closed for enrolment (cohort 2 since July 2018; cohort 1 since November 2018). OptiRif Kids will pave the way for the design and evaluation of shorter course regimens for DS-TB in children using higher doses of rifampicin. The third and final cohort opened for enrolment in Q4 of 2018, and completion of this cohort and closing of the study, with final analyses, is expected in 2019.

# Therapeutic trials: prevention of MDR-TB

TB-CHAMP (MDR-TB preventive therapy trial): (DTTC PI: Hesseling, Schaaf, Seddon) This is the first ever randomized phase III placebo-controlled trial to assess the efficacy of levofloxacin preventive therapy vs. placebo in child contacts of multidrug-resistant (MDR) TB. The sample size is approximately 1500 children 0-5 years of age. The trial, led by DTTC, is funded by the BMRC/Wellcome Trust/DFID and includes 3 South African sites, including Shandukani (WHRI, Johannesburg, Dr. Lee Fairlie), PHRU Matlosana, Klerksdorp (Prof. Neil Martinson) and DTTC. A formal PK lead-in study was completed in 2017 looking at bio-availability of a novel scored dispersible levofloxacin formulation (to be used in the main trial) in 24 children. This study showed that the paediatric levofloxacin formulation was well tolerated, acceptable, safe, and had adequate exposures in children < 5 years of age. Pharmacokinetic and acceptability data from this study have been recently published in two journal articles. The main trial opened to accrual in Q4 2017 at DTTC, and in Q1 2018 at the other 2 sites. As of 31 January 2019, 393 children had been screened and 3015 enrolled in the study. Additional grant funding has been requested from the The European & Developing Countries Clinical Trials Partnership (EDCTP) and other funders. This trial is likely to have a considerable impact on global and national guidelines and on clinical care.

# Therapeutic trials: treatment of MDR-TB

**Otuska 232/233:** (PI Anthony Garcia-Prats): Along with a site in the Philippines, the DTTC is implementing this industry sponsor-funded (Otuska, Japan) Phase 1 (232) and Phase 2 (233) trials which seek to characterize the pharmacokinetics and safety of delamanid in children with MDR-TB. New drugs are desperately needed for the treatment of children with MDR-TB, and these trials are critical for ensuring timely access to this important new medication. In this age de-escalation trials, all age groups (0 to 17 years) have fully enrolled for both studies. Data from these groups informed WHO guidance in 2016 for the use of delamanid in children 6-17 years of age with MDR-TB, and in 2018 for Group 3 (ages 3-6 years). The 232 study is now closed with final data analysis ongoing, and

long-term follow-up of Group 4 (0 to 2 years) completed at the end of 2018, with the final analysis now ongoing.

MDR PK 1: (PI Anneke Hesseling) This NIH-funded (R01 grant) was completed during 2016. This study of the pharmacokinetics and safety of routine doses of existing second-line anti-TB drugs in HIV-infected and uninfected children, was the first study of its kind, in some cases generating some of the only data on the pharmacokinetics and safety of these medications in children with TB. Over 4 years 312 children aged 0-15 years were enrolled and followed long-term for safety and treatment outcome. This study has already resulted in seminal data on the use of levofloxacin, ofloxacin, moxifloxacin, amikacin, high dose isoniazid, and the effect of MDR-TB treatment on the pharmacokinetics of ARVs commonly used in children. Analysis of para-aminosalicylic acid (PAS), ethionamide, terizidone, clofazimine and linezolid are ongoing. Long-term outcome and toxicity data are being analyzed currently. These data are informing international guidance on the dosing of these medications in children. The platform generated from this study has supported 5 PhD students and 6 master's degree students, to date.

MDR PK 2: (PI Anthony Garcia-Prats) Building on the data, experience, and clinical platform of MDR PK1, this NIH-funded (R01 grant) seeks to evaluate the pharmacokinetics and safety of modeloptimized doses of the key second-line anti-TB medications levofloxacin, moxifloxacin, and linezolid in children treated for MDR-TB. The study will also characterize the acceptability and palatability of different dosing strategies and will evaluate the effect of formulation manipulation on pharmacokinetics in children. The total sample size is n= 100; to date, 84 children have been enrolled through the end of 2018, and accrual is now closed with long-term follow-up expected to be completed in 2019. Data from the study on linezolid and moxifloxacin pharmacokinetics was shared with the World Health Organisation (WHO) in 2018 and informed revised dosing guidance for these medications. Multiple analyses on the pharmacokinetics, safety and acceptability of linezolid, moxifloxacin and levofloxacin, as well as analysis of treatment outcomes are planned in 2019. **IMPAACT P1108**: ( PIs Anneke Hesseling, Simon Schaaf) This NIH-funded IMPAACT network phase I/II trial to determine the optimal and safe dose of bedaquiline in HIV-infected and uninfected children with MDR-TB has been in development for 4 years. The protocol was completely developed and received MCC approval in August 2016. Five international sites, including DTTC, 2 other South African sites, one in India and one in Haiti, opened up to accrual on the 1st of September 2017. Bedaquiline has become a critically important treatment option for adults with MDR-TB, with extensive roll-out of the drug within the routine TB programme setting in South Africa and many other countries; however the lack of data in children has prevented paediatric access. This trial will provide desperately needed data on bedaquiline in both HIV-infected and –uninfected children with MDR-TB in order to ensure paediatric access to this new TB medication. The first participant was enrolled on the 21 September 2017. The first 15 children have been enrolled and the first formal interim analysis has been completed. Interim data was shared with the WHO and informed new dosing recommendations for bedaquiline in children. It is expected that the two youngest cohorts (0 to 2 years, 2 to 5 years) will open in early 2019.

Bedaquiline CRUSH Study (TASK-002): (Protocol Co-Chair – Anthony Garcia-Prats). This study, funded by National Institute of Child Health and Human Development (NICHD) through the IMPAACT network was a randomized, open label, crossover, bioequivalence study to assess the bioavailability of bedaquiline given as whole tablets or suspended (dissolved) in water. The study was conceived in order to facilitate the use of the bedaquiline whole tablets for use in paediatric clinical trials and eventually in routine care, as the paediatric formulation is not available now or expected to become available for some time. Twenty-four healthy male and female volunteers were randomly assigned 1:1 to one of 2 treatment sequences in order to receive either first a single dose of the crushed form of bedaquiline, as the experimental, and secondly a single dose of the whole tablet as the approved dosing form, or vice versa. The study was implemented at TASK Applied Science in Cape Town, and analysis completed in 2017 (E Svensson, Upssala University). Results, which showed that bedaquiline administered dissolved in water was bioequivalent to whole tablets, were published in

2018 and will inform the practical use of the adult formulation in children in the field, where access to the paediatric formulation is expected to be limited for some time.

# Lung health, diagnostic and biomarker studies

(PIs Elisabetta Walters, Marieke van der Zalm and Anne-Marie Demers)

The diagnostic platform nested in the DTTC paediatric programme focuses on improving the diagnosis of TB in children. TB in children is mostly clinically diagnosed as the collection of high-quality sputum samples is resource-intensive and relatively invasive and available laboratory methods are insufficiently sensitive to detect the low organism concentration typically found in samples from children. However, young children are at risk of delayed diagnosis due to poor diagnostic tools, resulting in increased risk of morbidity and mortality from advanced TB. The overall aim of the diagnostic platform is to improve the detection of TB in children using comprehensive strategies that are feasible, child-friendly and adequately sensitive to detect paucibacillary disease, focusing especially on young children.

**UMOYA** ("breathe"). Intra-thoracic tuberculosis in children: moving towards better diagnosis and improved lung health: (PIs Liz Walters, Marieke van der Zalm). A diagnostic cohort study that builds on a previous diagnostic study led by Liz Walters and has leveraged funding from NIH, Thrasher Foundation, EDCTP and the South African MRC. UMOYA started recruitment November 2017 and currently has 102 children enrolled. The aim is to enroll 300 children with suspected TB and 100 healthy sibling controls from Tygerberg hospital and Karl Bremer hospital. The study will continue to evaluate improved diagnostic strategies for paediatric TB, including novel laboratory techniques for both molecular and culture-based diagnosis on respiratory and stool specimens. It will support ongoing evaluation of blood and urine biomarker work. In addition, the study will continue work to evaluate the interaction between viral and bacterial co-pathogens with TB, and plans to assess and monitor lung function longitudinally in all children

#### **Evaluation of novel TB biomarkers in children**

**Study 1. Urine proteomics:** This project aims to identify host-derived urine proteins that are differentially expressed in children with TB compared to children with non-TB illness. The project is in collaboration with US partners Robert Husson and Hanno Steen from Boston Children's Hospital, USA. This year, the focus of the project has been to optimise the LC-MS/MS platform to achieve a high detection of proteins in urine samples. This is done in the Steen laboratory at Boston Children's Hospital, Harvard University. Data analysis will include both traditional logistic regression approaches (lead by Harvard University) as well as bio-informatics methods (lead jointly by Harvard and Stellenbosch Universities). A doctoral student, Ashley Ehlers, under the supervision of Prof D Tabb from SU's Division of Molecular Biology and Human Genetics, has been awarded a scholarship by Boston Children's Hospital, to work on this project.

**Study 2. Serum bio-signatures**. This project aims to identify pathogen-derived serum antigen peptides that are differentially expressed in children with TB compared to children with non-TB illness, using a novel NanoShell-MS assay. The project is in collaboration with US partner Tony Hu from Arizona State University.

**Study 3. TB lipoprotein biomarkers**. This exploratory study aims to identify whether TB lipoprotein is a useful diagnostic biomarker of active TB disease in children. Results of an initial analysis are in progress. TB Lipoprotein (TLP) is a specific mycobacterial modification of low density lipoprotein (LDL), a host biomolecule.

# **Epidemiological and operational research for pediatric TB**

Please refer to <u>Research focus Area Two: Health Systems and Operational Research</u> for updates of pediatric focused epidemiological and operational research studies.

# Research focus Area Two: Health Systems and Operational Research

The aim of the research in this pillar is to help improve TB and HIV care by building an evidence base for effective programme implementation. Operational research (OR) focuses on identifying gaps in health programme quality, efficiency and effectiveness, evaluating factors that contribute to these, and testing interventions to improve outputs and outcomes.

This research focus area also include epidemiological and routine data analyses, modelling, and health system strengthening research, all contributing to the overall aim of generating evidence to strengthen program implementation. Outputs for 2018 are listed under the following headings: Health systems Research and Operational Research (including TB mortality and childhood TB as specialized focused areas).

# **Health Systems and Operational Research**

LINKEDin: reducing initial loss to follow up among tuberculosis patients: (PIs: Anneke Hesseling and Muhammad Osman). This study is funded by the Bill and Melinda Gates Foundation and will be implemented in 2019. Individuals diagnosed with tuberculosis (TB), but who are lost to follow up before initiation of TB treatment and/or TB notification are referred to as "initial loss to follow up" (ILTFU). This study aims to: 1) demonstrate the impact of an automated notification system of TB patients in hospitals on increasing the number of notified TB cases; and 2) demonstrate the impact of the implementation of an automated alert-and-response TB patient management system on increasing the proportion of TB patients who are linked to TB services and who initiate treatment at different levels of health care. The study will be implemented in 6 hospitals and 31 primary health care facilities in 3 provinces of South Africa; Western Cape (WC), Gauteng (GP) and Kwa-Zulu Natal (KZN). The study will include all laboratory-diagnosed TB patients in the identified study sites as well as clinically diagnosed TB patients in the WC. Using a before/after design, the study will measure the proportion of TB patients who are ILTFU at baseline and again after the 12th month, following the implementation of the study interventions. Between July and December 2018, the study received ethics approval with the Stellenbosch University Health Research Ethics Committee (HREC) and approvals from the relevant health departments/authorities. Numerous stakeholder engagement meetings were held to introduce the study and get buy-in at all levels. Interviews were held for key staff. A baseline data collection tool was designed.

# **Operational Research**

Evaluating 10 years of TB control in South Africa: (PIs: A.C. Hesseling/Muhammad Osman). Routine TB surveillance data have been under-utilised in identifying and understanding programme weaknesses and finding ways to address these. The socio-economic determinants that may be contributing to poor TB control have not been adequately evaluated. As South Africa scales up TB control efforts to achieve the even more ambitious "End TB Strategy" goals, a more data-driven approach to TB control efforts is required. This study aims to undertake a comprehensive, in-depth analysis of TB control in South Africa to review epidemiological trends from 2004 to 2018 and to identify achievements and programme gaps where intervention is required. This project is not funded and is being implemented in collaboration with National Department of Health. Abstracts presented at the 49<sup>th</sup> Union World Conference on Lung Health in The Hague in October 2018

1. Trends in the South African tuberculosis epidemic with scale up of TB-HIV integration services (poster) (Muhammad Osman, Karen du Preez, Rory Dunbar, Alex Welte, Pren Naidoo, Anneke C. Hesseling). This poster highlighted HIV testing and ART coverage in SA have improved substantially over 10 years. The burden of TB amongst HIV-infected individuals is declining but remains substantially greater than among HIV uninfected. The overall declining trend in the TB epidemic appears less significant among the HIV-uninfected population.

- 2. Mortality amongst HIV-positive female tuberculosis patients in South Africa (short oral) (MM Claassens, C van Schalkwyk, R Dunbar, M Osman, P Naidoo, M Borgdorff). This short oral presentation focused on data between 2009 and 2013. It demonstrated a substantial decline of mortality in HIV-positive TB patients, while among HIV-negative TB patients, mortality was stable. Limitation regarding the time of ART initiation, CD4 count or ART adherence were recognized and updated analyses including data through to 2017 is in progress.
- 3. Ten years of routine paediatric TB surveillance data in South Africa high disease burden and poor treatment outcomes (oral) (K Du Preez, M Osman, P Naidoo, S Dlamini, AC Hesseling). This oral presentation showed the significant number of children treated for TB and reported in ETR.net in South Africa. The proportion of children overall (11-13%), and proportion of these <5 years of age (62%) are consistent with that expected in high burden TB settings. Documentation of HIV testing has improved over time but the pre-treatment microbiological confirmation of TB disease remains low (9%). Ongoing analysis is exploring age, TB disease spectrum, HIV co-infection and treatment and reasons for poor treatment outcomes in children.</p>

**Operational Research Assistance Program:** The overall aim of ORAP is to undertake operational research as an integral component of health programmes in South Africa to contribute to improved quality and performance of the health system.

In November 2016, a new miniature version of ORAP was implemented in the Western Cape. Participants (5 from health services) have embarked on an experiential learning course to develop 3 independent study protocols for research to be undertaken. Five mentors from DTTC are supporting the trainees through protocol development, study implementation and publication of their findings. All 3 projects have completed the mentored protocol development stage and were approved for implementation by the Stellenbosch University HREC.

- Comparing early treatment outcomes in multidrug resistant (MDR) tuberculosis (TB) patients treated with conventional MDR-TB regimens is being implemented by Dr L Naidoo and Dr D Da Costa and supervised by M Osman and A Garcia-Prats. Data collection and capture is complete and 177 records are in the process of data cleaning before analysis. Abstracts were presented at the South African TB Conference in Durban in June 2018 and the 49<sup>th</sup> Union World Conference on Lung Health in The Hague in October 2018.
- Identifying risk factors associated with unfavourable treatment outcomes amongst patients treated for Isoniazid mono-resistant TB in Cape Town, South Africa is being implemented by Dr K Joseph and Dr F Solomon-Da Costa supervised by K du Preez. Data has been received on 370 patients and preliminary analysis is in progress. An abstract was presented at the 49<sup>th</sup> Union World Conference on Lung Health in The Hague in October 2018.
- 3. Motivations For, Anxieties And Preferences (MAP) About Transitioning To Community-Based Art Adherence Clubs (CAC) Among Patients Attending Facility-Based Art Adherence Clubs is being implemented by Dr R Mabezue and supervised by R Dunbar and M Osman. Data collection is in progress and over 200 participants have been consented and interviewed.

# Operational Research focused on TB mortality

Former TB patients as a high-risk group for TB-associated morbidity and mortality in high-incidence communities in Cape Town − a pilot study (PIs: Florian Marx and Muhammad Osman). A high risk of TB, chronic lung disease and mortality have been reported among people with a history of previous TB treatment, but data from high-incidence settings remain limited. This study aims to locate adults who had successfully completed TB treatment in the past five years, and to characterize general morbidity and mortality. Routine electronic TB treatment register data was used to randomly select 256 adults (≥18 years) who successfully completed pulmonary TB treatment between 2013 and 2017. Household visits were conducted in a high-incidence community in suburban Cape Town to locate and survey former TB patients. Individuals who could produce sputum were bacteriologically

investigated for TB. Household interviews and vital registration data were used to estimate mortality among patients who could not be traced.

We have interviewed 51 former TB patients and 52% (19/36) were living with HIV. Approximately half had self-reported symptoms indicative of chronic lung disease and of 28 participants who provided a sputum sample, one (3.6%) had culture-positive TB and another two reported currently being on re-treated for TB treatment, providing 3/51 prevalent cases of TB. Among 256 former TB patients, there were 15 (5.9%) confirmed deaths were confirmed (5.9%) with a median time of 1.43 (IQR: 0.48-2.16) years between the successful completion of the previous TB treatment episode, and death. The mortality rate among former TB patients was 2.7 deaths (95% CI: 2.5-3.0) per 100 person years with a standardized mortality ratio of 4.1 (95% CI: 3.6-4.7) compared to the general local population.

Our preliminary results are consistent with a high prevalence of respiratory disease including prevalent TB among former TB patients, and higher mortality.

The abstract: Finding former tuberculosis patients - a high-risk group for tuberculosis-associated morbidity and mortality in Cape Town, South Africa: A pilot study, was published as part of SA MRC 12<sup>th</sup> annual Early Career Scientist Conference and presented as a poster at the SACEMA Research days 2018.

This study was funded through a research grant (R64 000) provided by the Günther Labes Foundation and the Oskar Helene Heim Foundation (Berlin, Germany) to FMM and an Emerging Research Award (R35 000) provided by DST-NRF South Africa Centre of Excellence in Epidemiological Modelling and Analysis (SACEMA), Stellenbosch University to MO.

Retrospective descriptive study of sudden and unexpected deaths related to TB in Cape Town: (PI: Muhammad Osman). Undiagnosed TB as a final or contributing cause of death is difficult to quantify but has been estimated through post mortem studies. In a low TB incidence setting (Ireland), only 15 cases of TB were diagnosed in a series of 4930 autopsies while in India, a high TB burden setting, 8.7% of autopsy cases were diagnosed with active tuberculosis and 60% of these were not diagnosed before death. The under ascertainment of both TB and TB mortality in a high burden TB/HIV setting warrants further evaluation. We aimed to estimate TB disease as a final or contributory cause of death among sudden unexpected death in adults and children in Cape Town. We have undertaken a collaborative project with SU Forensic Pathology Services (FPS) and have retrieved and captured data on 719 cases of sudden and unexpected death in 2016. 79 records are expected to be complete in Q1 2019. This study has been supported by Prof Anneke Hesseling through SA NRF SARChI funding.

# Operational research focused on childhood TB

The aims of this research area are to firstly evaluate and identify gaps in the current health systems preventing and providing care to children with TB, and secondly to pilot and measure the impact of innovative solutions through rigorous implementation science methodology.

This research area is led by Dr Karen du Preez, and current projects focusses on optimizing pediatric TB surveillance strategies and strengthening prevention and routine data for children with TB, including HIV co-infected children.

# Hospital focused surveillance strategies for childhood TB

The results of three studies related to hospital surveillance for childhood TB were published in 2018 – two focused on TB surveillance at tertiary hospital level, specifically children with TB meningitis (TBM) and spinal TB, and the other focused on TB surveillance at a district level hospital.

1. The impact of BCG shortages on the burden of childhood TBM (*Published in The Lancet Global Health, January 2019, see Appendix I*)

During 2017, we have observed a substantial increase in the number of children admitted for diagnosis or management of TBM or tuberculomas to the paediatric neurology ward at Tyberberg Hospital. Birth cohort analyses of admitted children <2 years of age with TBM/tuberculomas found

nearly three times as many children born in 2015 compared to the mean number of children born in 2012 to 2014. We hypothesize that this increase is related to the global BCG shortages, which has impacted substantially on the number of BCG available during 2015 in the Western Cape Province.

2. The burden and trend of spinal TB in children (*Published Epidemiology and Infection, 2018, see Appendix I*)

This retrospective study, led by Dr Theresa Mann from the Division of Orthopedic Surgery, assessed the overall burden and trend in spinal TB at tertiary hospitals in the Western Cape Province during 2012-2015. A total of 74 children and 319 adults were identified. The study found a substantial burden of spinal TB cases with a high proportion of children presenting with severe disease and requiring corrective spinal surgery. Although spinal TB affects a relatively small proportion of TB cases, the burden on public health services remains substantial in high TB burden settings, and earlier diagnosis particularly for child cases are needed.

3. The burden, spectrum and outcomes of childhood TB disease at a district level hospital (*Published IJTLD, 2018. See Appendix I*)

This retrospective cohort study aimed to characterize the burden, spectrum and TB treatment outcomes of children managed at a district level hospital in Cape Town. Nearly 100 children were managed at the district level hospital over a 7 month period in 2014. The majority (85%) were very young (≤2 years), a third (34%) were malnourished and a fifth (19%) had HIV co-infection. Although 96% of children successfully continued care after hospital discharge, favorable TB treatment outcomes (cured/treatment completed) were only recorded for 78%. This study found a substantial burden of paediatric TB are managed at district hospital-level, with patients often being very young and frequently presenting with comorbid diseases. TB prevention opportunities and treatment outcomes could be further improved.

#### Routine TB treatment surveillance at national level:

Within the study, "Evaluating 10 years of TB control in South Africa", multiple analyses have been completed and are planned to specifically look at the TB epidemic in children in South Africa over time. One abstract was accepted for an oral presentation at the 49<sup>th</sup> Union World Conference on Lung Health in 2018: Ten years of routine paediatric TB surveillance data in South Africa - high disease burden and poor treatment outcomes (oral) (K Du Preez, M Osman, P Naidoo, S Dlamini, AC Hesseling). Main findings of the abstract have been detailed earlier in this section.

#### Research Focus Area Three: HIV Prevention Research

The DTTC has made great strides in the field of HIV prevention with two studies as part of the HIV Prevention Trials Network (HPTN) conducted during 2018. While the HPTN 071 (PopART) study concluded in 2018, the HPTN 084 (Life) was initiated early in the year.

HPTN 071 (PopART): (Lead: Nulda Beyers, Peter Bock). The HPTN 071 or PopART study is a cluster-randomized trial which aimed to determine the impact of two community-level combination prevention packages, both of which included universal HIV testing and intensified provision of HIV/ART care, on population-level HIV incidence. The study was conducted in 9 communities in South Africa and 12 in Zambia. Communities were randomized to arms A (full intervention prevention package plus ART regardless CD4 count), B (full intervention prevention package plus ART according to government guidelines) or C (standard of care). Following changes to ART guidelines during the study, Arm A and B both offered ART regardless of CD4 count from Oct 2016 onwards. Despite this change the study was still well powered to evaluate the primary outcome, HIV incidence. At each of the arm A and B sites, interventions were delivered to the entire community by a cadre of Community HIV-care providers (CHiPs) with referrals to government Primary Health Care (PHC) clinics. CHiPs provided clients with condoms, screen and refer relevant clients to government clinics for HIV, TB and STI treatment and voluntary male medical circumcision (VMMC). Primary outcomes were measured in a randomly selected individual level cohort of approximately 2000 individuals in

each community (including the Arm C communities) over 36 months referred to as the population cohort (PC).

Field work for PopART began in January 2014 and finished in June 2018. For the CHiPs intervention, by the end of 2017 152,454 households had been visited in South Africa and Zambia and 168,556 individuals tested for HIV. At the end of PC follow up, in June 2018, more than 38 000 participants had completed the final round of the PC follow up PC36. The PopART data were revealed in Seattle in December 2018 where the team completed the primary analysis comparing HIV incidence across study arms. The primary results will be presented at CROI in March 2019. Dissemination of primary results to key stakeholders including study communities will be completed in March 2019.

**HPTN 084 (Life) trial:** (PI: Dr PA Bock) The HPTN 084: A Phase 3 Double Blind Safety and Efficacy Study evaluating Long-Acting Injectable Cabotegravir Compared to Daily Oral TDF/FTC for Pre-Exposure Prophylaxis in HIV-Uninfected Women. The primary end point is HIV incidence. Participants will be followed up for 3 to 4 years. Overall the study aims to enroll 3,200 participants across the 20 sites (approximately 160 participants per site).

The Kuilsriver site office was newly established from January 2018. DAIDS site assessment visit was completed in March 2018, the DAIDS laboratory assessment in April 2018 and the site was activated by DAIDS & HPTN by August 2018. The team enrolled their first participant in September 2018 and by end December 2018 had enrolled a total of 16 participants. The team also successfully completed a further audit in December 2018. The focus in the first two quarters of 2019 will be scale up community activities and recruitment.

#### **Crosscutting Research (Supporting all three research focus areas)**

#### **Sociobehavioural Sciences**

The Sociobehavioural science team have successfully implemented a broad range of data collection and analysis activities as components of the wider DTTC research portfolio and contributed to all three research focus areas. These activities ranged in scope from innovative ways to assess children's experiences of TB treatment palatability (Project Masterchef) to an exploratory analysis of the potential of technology for TB treatment adherence support (TB PAKS). Across activities, all data collection, management, processing and preparation for analysis is standardised and managed by a dedicated data quality team. In 2018, the team transcribed approximately 780 hours of recorded interview/discussion data and translated over 3000 pages of these transcripts from Xhosa or Afrikaans to English. Further, the team again proved to be highly innovative in designing and refining novel methods to answer research questions with greater sophistication. A priority for 2019/2020 is to support staff in post-graduate studies, and in doing so, building on the culture of academic learning and publication.

# Sociobehavioural Science and Paediatric TB

**SHINE**: In-depth acceptability interviews and observations with caregivers and children about a novel, child-friendly formulation of a fixed-dose combination of first-line TB treatment to be used in the SHINE trial. The SHINE trial offers participants a fixed-dose combination child-friendly formulation of first-line TB treatment. This set of 20 days of semi-structured observations and discussions with 16 caregiver/child dyads informs the acceptability of this formulation for use and clarifies potential misunderstandings by participants about administration of the treatment on trial. A manuscript reporting findings has been developed with submission for publication anticipated in early 2019.

**TB-CHAMP:** In-depth acceptability interviews and observations with caregivers and children about novel formulation of levofloxacin to be used in the TB-CHAMP trial: The TB-CHAMP trial uses a novel, child-friendly formulation of levofloxacin. This set of 30 days of semi-structured observations and discussions with 17 caregiver/child dyads informs the acceptability of this formulation for use and clarifies potential misunderstandings by participants about administration of the treatment on trial.

Data processing is ongoing, and a manuscript is in preparation for submission in 2019 as part of Dillon Wademan's PhD.

**Impact of MDR-TB on children:** Exploring caregivers' and health workers' perceptions on the effects of caregiver-child separation during long-term hospitalisation for MDR-TB in the Western Cape: A qualitative study. This qualitative project, conducted by Kyla Meyerson in completion of her MA in Psychology (SUN), addressed the third objective of MDR-PK II namely, to characterise the acceptability of DR-TB treatment. MDR-TB treatment requires extended hospitalisation which currently entails caregiver-child separation. Caregiver-child separation has been shown to cause behavioural and emotional problems in children. We explored caregivers' and health workers' perceptions of the effects of caregiver-child separation during long-term hospitalisation for MDR-TB treatment. We conducted 19 semi-structured, in-depth interviews with caregivers and health workers of children (aged zero to five years) who were receiving hospital-based treatment for MDR-TB. A thematic analysis was conducted to organise and interpret the data. Three major themes were identified: (i) MDR-TB treatment was a traumatic experience; (ii) Children's behavioural and emotional states during MDR-TB treatment included excessive crying, aggression, hyperactivity, and withdrawal; (iii) Caregivers' and health workers' behavioural and emotional management strategies included deception, threat, and prioritisation of biomedical health over psychological health. Children's, caregivers', and health workers' problems that emerged during MDR-TB treatment most likely result from a complex interplay between various factors; including precipitating factors of MDR-TB treatment and caregiver-child separation, and predisposing factors of poverty and exposure to violence.

MDR-PK - Masterchef: To understand children's and caregivers' experiences of acceptability (including palatability) of the key second-line tuberculosis drugs, within the MDR-PK II project. Over a four-week period, we conducted interviews with 16 paediatric participants (and where appropriate their caregivers). Our research consisted of a semi-structured interview schedule and various participatory research activities. In the latter, we asked children to rate various tastes, forms of medicine (for example, tablet, capsule, syrup, chewable pill and injection) and colours and forms of medicine packaging in an attempt to design the ideal MDR-TB medication. We wrote a report on the preliminary findings and disseminated these preliminary findings to DTTC health workers at Brooklyn Chest Hospital. Data has been processed to prepare for an in-depth analysis.

Lung health in Africa across the life course (LuLi): The purpose of this project is to describe the prevalence and clinical characteristics of post-tuberculosis lung disease among former tuberculosis patients in Cape Town, South Africa. The project utilised a multi-disciplinary approach, forming a collaboration between the Departments of Physiotherapy, Pulmonology and Paediatrics and Childhealth (DTTC). Our team at DTTC were involved in the recruitment and implementation of the project in one community in the Western Cape of South Africa. A total of 45 participants were recruited to the project. Participants underwent several physiological and lung function tests, as well as completing a few psycho-social surveys to provide a holistic overview of the participants' health. The project aims to define the burden of symptoms and functional deficits in non-healthcare seeking patients who have completed TB treatment in order to build a platform to address potential interventions for those affected.

TB PAKS: Paediatric Adherence, Knowledge and Support: The TB PAKS project aims to develop and refine an 'app' suitable for smartphones to be used by children (and their caregivers) who are on TB prophylaxis or treatment in order to improve their experience by supporting learning about TB, provide adherence reminders/virtual rewards, and facilitate management of the social impact of TB such as disclosure. Preliminary work on the as-of-yet unfunded project began in 2018, with the Sociobehavioural team hosting two students whose work is helping inform development of the app. Ava Archey, high-school student and daughter of HIV/TB researcher Kelly Dooley (honourary associate professor, Department of Paediatrics and Child Health), visited the DTTC for two weeks during her summer break to develop a prototype of a TB mobile application (app) for children following a literature review of TB disease and technology. She presented the prototype to

stakeholders at the DTTC. Rachel Morse, an undergraduate student from King's College London, interned at the Centre for two months as part of a larger collaboration between Stellenbosch's Faculty of Medicine and Health Sciences and their Department of Global Health and Social Medicine. In her dissertation, which drew on qualitative data and observations collected across various TB projects at the Centre, she explored factors that influence children's adherence to TB treatment and considered ways in which a mobile 'app' could be used to address them. Both these outputs are providing an important foundation for the project.

TB-TREATS: Tuberculosis Reduction through expanded antiretroviral therapy and TB screening: The overall aim of this project is to measure the impact of a combined TB/HIV intervention of population level screening for TB, combined with universal testing and treatment (UTT) for HIV on TB incidence, prevalence and incidence of TB infection. The study is a collaboration between Zambart in Zambia, Health Systems Trust in South Africa, the London School of Hygiene and Tropical Medicine, Imperial College and the University of Sheffield in the UK and KNCV in the Netherlands. Given that the TREATS study is being conducted in the same communities as the HPTN 071 (PopART) in South Africa and Zambia, the Sociobehavioural Sciences Team at the Desmond Tutu TB Centre was contracted by the Health Systems Trust to conduct nested qualitative research as part of the formative work towards the TB-TREATS study. The first part of this research was conducted between October and December 2018. We conducted four in-depth interviews with key TB staff to describe their experience with TB service delivery and the role of other stakeholders (n=1 in each arm A and C site). Further research with TB patients is underway and will contribute to a broader understanding of the impact of TB in people's lives in South Africa.

#### <u>Sociobehavioural Science and HIV Prevention Research</u>

HPTN 071a (Stigma Ancillary Study) - Health Worker Open Cohort Survey: The Stigma Ancillary Study was a sub-study to the PopART (HPTN071) trial. The stigma study is a mixed method study that nests enquiry about HIV stigma across the PopART study. In particular, it the study examines how health care worker knowledge, attitudes, and experiences of delivering HIV care are influenced by, and influence, the delivery and uptake of the PopART intervention. The quantitative component of the study involved a self-administered questionnaire offered to all health care workers (including CHiPs) working in PopART study communities. Data collection was done in the three waves in 2014, 2015 and 2017. By June 2017, data collection had been completed in South Africa with 797 health workers enumerated. In 2018, the stigma activities included cleaning data from all three rounds and preliminary analysis conducted for health facility dissemination activities. Several manuscripts are in preparation.

HPTN 071 PopART for Young People (P-ART-Y) Study: Uptake and acceptability of a combination HIV prevention package among young people in Zambia and South Africa. The P-ART-Y study concluded in March, 2018. The primary outcome of the study was the uptake of voluntary HIV counselling and testing, within the previous 12 months at the time of survey administration, amongst adolescents aged 15-19 years. This served as a comparator for the primary outcome of the study (knowledge of HIV status). Other outcomes included acceptance of HIV testing and retesting. Linkage to care, antiretroviral therapy (ART) adherence and retention in care, uptake of prevention to mother and child transmission (PMTCT), voluntary medical male circumcision (VMMC) and condoms. Qualitatively, we looked at the acceptability of the PopART interventions and barriers to access in adolescents. We also looked at the experiences of adolescents living with HIV. The particular needs and vulnerabilities of adolescent girls, including those living with HIV, were also studied. Data collection was completed on 4 November 2017. Data cleaning was completed in 2017 and analysis is ongoing with three manuscripts in development for submission in 2018.

**P-ART-Y Dissemination:** The study team is worked alongside the Adolescent Advisory Board (ACAB) and Community Engagement Team on the planning of dissemination events. These took place in the Arm C site of the HPTN 071 (PopART) trial and were conducted on the 9, 16 and 23rd March and the 13<sup>th</sup> of April 2018. Preliminary findings of the cross-sectional survey were shared with young people

some of whom participated in the study. These findings included survey uptake frequencies by age and gender and some of the secondary objectives. The primary outcome findings will be disseminated in 2019 together with results from the main trial, the HPTN 071 (PopART) trial. The dissemination events were well received by young people and this forum served as a platform for further discussion on the findings.

Mixed-data in-depth interviews with staff/stakeholders and operational review of clinic-based components of the HPTN 071 (PopART) intervention package: The PopART intervention package included a large number of health system innovations and implementation lessons from the health-facility delivery of universal testing and treatment. Specific focus areas were (a) preparation for transition to treatment regardless of CD4-count, (b) health-facility based HIV testing services, (c) explaining ART regardless of CD4-count to clients, and (d) implementing adherence support structures. Between February and April, 26 in- depth interviews with key informants were collected. Mixed data observational and discussions with field staff of the community-based components of the HPTN 071 (PopART) intervention package – annual round 5: A set of structured observations and discussions with implementers conducted annually in 9 study communities between January and March. The final round of data were collected without issue in 2018.

HPTN 071 (PopART) Research Collective (PRC): PRC is a collaborative project lead by Dr P Bock, composed of DTTC PopART staff and PopART implementing partners (City of Cape Town, Western Cape DoH, ANOVA Health Institute, Kheth' Impilo and Cape Winelands DoH). This consortium is aimed at formulating lessons learned of implementing the HPTN 071 study at health facilities, over trial period. The PRC consists of five groups each with an aim of developing a manuscripts related to the implementation of the PopART intervention. Data collection was completed in November 2017. A total of 51 interviews were conducted with health services staff at a provincial, district, regional and clinic level. Manuscripts are under development.

Sociobehavioural Science Students Graduated

- 1. Rene Raad MA in Anthropology at SUN
- 2. Hermine Kruger Honours in Psychology at SUN
- 3. Arlene Marthinus Honours in Psychology at UWC
- 4. Msondezi Ketelo BA in Anthropology at UWC

#### **Epidemiological Modelling**

Modelling of tuberculosis and the impact of TB control interventions constitutes an additional core competency to be built at DTTC in the forthcoming years. In 2018, a DTTC Working Group Data Analysis and Modelling (WGDAM) was established to build and strengthen capacity for in-house modelling (see below). A strategic TB modeling partnership and research programme between DTTC and the South African Centre of Excellence in Epidemiological Modelling and Analysis (SACEMA) was prepared in 2018 and is set to take off in 2019. This novel initiative is intended to provide a platform for mathematical modelling and intervention research to inform decision making for TB control at national level in South Africa as well as internationally.

#### Working Group: Data Analysis & Modeling (WGDAM)

A DTTC Working Group Data Analysis and Modelling (WGDAM) was established to build and strengthen capacity in mathematical modeling of TB and related control interventions. The working group is led by Dr Florian Marx and holds weekly meetings at DTTC. It currently consists of 10 members with backgrounds in medicine, epidemiology/public health, data management and statistics. In 2018, WGDAM initiated a joint research project which aims to project the population-level impact of ART scale up on TB in incidence and mortality in the nine South African Provinces. Prior to this project, the group reviewed data from the National Antenatal Sentinel HIV Prevalence

Survey and the District Health Barometer to compare rates of TB and HIV in the 52 South African health districts. This study highlighted poor association between HIV prevalence estimates and rates of reported TB at subnational level, which has important implications for TB control in South Africa. Efforts are needed to better understand the determinants of local variation in TB and HIV rates in South Africa. Other modelling projects involving WGDAM members in 2018 include studies estimating the impact of TB household contact investigation and preventive therapy, of rolling out novel rapid diagnostic tests in South Africa, and of targeting TB case finding and prevention to previously treated people. The following manuscripts have been published:

- Sloot R, Maarman GJ, Osman M, Marx FM. Variation in HIV prevalence and the population-level effects of antiretroviral treatment for reducing tuberculosis incidence in South Africa. (Correspondence), on behalf of the Desmond Tutu TB Centre Working Group Data Analysis and Modelling (DTTC-WGDAM). South African Medical Journal 2018; 108(8): 606.
- 2. Dunbar R, Naidoo P, Beyers N, Langley I. Improving rifampicin-resistant tuberculosis diagnosis using Xpert((R)) MTB/RIF: modelling interventions and costs. Int J Tuberc Lung Dis 2018; 22(8): 890-8.
- 3. Marx FM, Yaesoubi R, Menzies NA, Salomon JA, Bilinski A, Cohen T. Tuberculosis control interventions targeted to previously treated people in a high-incidence setting: a modelling study. The Lancet Global Health 2018; 6(4): e426-e35.

The potential impact of household contact management on childhood tuberculosis: a mathematical modelling study (*Published Lancet Global Health*): This study evaluated the impact of carrying out household contact management strategies on the global burden of childhood TB disease and childhood TB mortality. By modelling the current situation for children following household exposure to infectious TB and assuming no household interventions, and then comparing it with two scenarios in which exposed children were screened for disease and then treated for TB infection, the impact of these interventions could be evaluated. Full implementation of household activities could result in a reduction of 160,000 cases of childhood TB and prevent 108,000 child deaths.

1. Dodd PJ, Yuen CM, Becerra MC, Revill P, Jenkins HE, Seddon JA. Potential effect of household contact management on childhood tuberculosis: a mathematical modelling study. The Lancet Global Health 2018; 6(12): e1329-e38.

#### **DTTC GRANT AWARDED: 2018**

| Description                                    | Funder                    | PI                          |
|------------------------------------------------|---------------------------|-----------------------------|
| LINKEDIN study (Reducing loss to follow        | Gates Foundation          | Anneke C. Hesseling/Muhamad |
| up among TB Patients)                          |                           | Osman                       |
| Consultancy for a systematic review of         | TB THINK TANK             | Tamaryn Nicholson &         |
| TB mortality in South Africa                   |                           | Muhammad Osman              |
| Consultancy paediatric MDR-TB                  | TB THINK TANK             | Jennifer Hughes             |
| National Guidelines                            |                           |                             |
| Protocol Vice Chair Funding                    | NIH IMPAACT network       | Jennifer Hughes             |
| Consultancy for a review of TB                 | TB THINK TANK             | Graeme Hoddinott            |
| preventative therapy implementation in         |                           |                             |
| South Africa                                   |                           |                             |
| HPTN 084 site                                  | NIH (HPTN)                | Peter Bock                  |
| 2 <sup>nd</sup> year extension of postdoctoral | Ramon Areces Postdoctoral | Elisa Lopez                 |

| fellowship                                        | Fellowship               |                                |
|---------------------------------------------------|--------------------------|--------------------------------|
| Travel fellowship Prof. Reza Yaesoubi at          | Stellenbosch University  | Florian Marx                   |
| the Yale School of Public Health, USA             | Scientific travel grant  |                                |
| Impact of viral co-infections on lung             | SA MRC                   | Marieke van der Zalm           |
| health in children with TB                        |                          |                                |
| Travel to the 22 <sup>nd</sup> International AIDS | Stellenbosch University  | Sue-Ann Meehan & Lario Viljoen |
| conference, Amsterdam in July 2018                | Scientific travel grant  |                                |
| BEAT Paediatric MDR-TB                            | UNITAID                  | Tony Garcia-Prats              |
| IMPAACT 2020                                      | IMPAACT NIH              | Tony Garcia-Prats              |
| SA MRC SHIP                                       | MRC grant for collateral | Anneke Hesseling, James Seddon |
|                                                   | funding for TB Champ     |                                |
| Graduate Internship Programme                     | NRF                      | Graeme Hoddinott & Anneke      |
|                                                   |                          | Hesseling                      |
| Graduate Internship Programme                     | NRF                      | Graeme Hoddinott & Anneke      |
|                                                   |                          | Hesseling                      |
| Familial management of paediatric co-             | Harry Crossley           | Dillon Wademan                 |
| morbid HIV, TB, and diabetes                      | (PhD scholarship)        |                                |
| Travel to the 49th Union World                    | Stellenbosch University  | Muhammad Osman                 |
| Conference on Lung Health, Amsterdam              | Scientific travel grant  |                                |
| in October 2018                                   |                          |                                |

#### **AWARDS AND OTHER RECOGNITION**

- 1. Professor Simon Schaaf received his 2<sup>nd</sup> A-rated NRF award in Port Elizabeth. September 2018.
- 2. Dr. Sue Purchase received the Discovery Award at the Durban TB Conference for overall "The Best Research Paper". July 2018
- 3. Dr. Mareli Claassens was appointed as a Research Fellow for DTTC.
- 4. Dr. Angela Dramowski was promoted to Associate Professor.
- 5. Professor Nulda Beyers received the Research Development Award. Prof Beyers mentored four successful PhD Graduates
- 6. Dr. Jana Winckler secured joint 3<sup>rd</sup> Place in the Young Scientist Award for podium presentation in the category basic and clinical pharmacology which included lab scientist and clinicians. 08 October 2018.

#### **DTTC ACADEMIC MEETINGS**

The TB Clinical Forum 2018, hosted by DTTC at the Stellenbosch University FMHS and organized by the City of Cape Town officials was held monthly between February and November 2018 with the objective of creating an interactive platform for academic researchers and government official in health services. Experts from the FMHS and City presented a broad range of relevant scientific research topics. An average of 50 participants attended these lively CPD- accredited interactional meetings on Friday afternoons, facilitating dialogue between service providers for TB and HIV care, and researchers at DTTC and the medical faculty. Four speakers from DTTC presented during 2018.

# **TB Clinical Forum 2018**

| DATE          | PRESENTER                    | TOPIC                                                                     |
|---------------|------------------------------|---------------------------------------------------------------------------|
| 9 Feb. 2018   | Dr. Lenny Naidoo (CCT)       | Highlights from the 48 <sup>th</sup> Union World                          |
|               | Dr. Jennifer Hughes (DTTC)   | Conference on Lung Health, Guadalajara,                                   |
|               |                              | Mexico                                                                    |
| 9 Mar. 2018   | Prof Marc Nichol –(NHLS/UCT) | Interpreting Ultra- A clinician's guide                                   |
|               |                              | Rapid diagnostic test & improved diagnosis                                |
| 13 Apr. 2018  | Prof. Helen Cox (UCT)        | What do we need to do to eliminate drug-<br>resistant TB???               |
| 11 May 2018   | Prof. Gary Maartens(UCT)     | Update on drug interactions between ARVs                                  |
|               |                              | and TB drugs                                                              |
| 8 Jun. 2018   | Prof. A. Diacon (TASK)       | Update on Adult MDR-TB treatment Trials                                   |
|               | Dr. Anja Reuter (MSF)        |                                                                           |
| 13 Jul. 2018  | ChloëShain (UCT)             | The Social Markers of TB                                                  |
| 24 Aug. 2018  | Prof. Graeme Meintjies(UCT)  | TB Lymphadenitis: Clinical management                                     |
| 21 Sept. 2018 | Dr. Tony Garcia-Pratts       | Ethics/Debate: "Do we really need the                                     |
|               | (DTTC)                       | injectable for managing DRTB?"                                            |
|               | D : ) ( A4 - d : d : (A455)  |                                                                           |
|               | Dr V. Mudalay (MSF)          |                                                                           |
|               | Dr A. Reuter (MSF)           |                                                                           |
|               | Dr J. Te Riele(BCH/WCG)      |                                                                           |
|               | br s. re niele(berly wed)    |                                                                           |
| 12 Oct.2018   | Prof. Anneke Hesseling       | DTTC-Update on Paediatric TB treatment Trials                             |
|               | Dr. Tony Garcia-Pratts       |                                                                           |
|               | (DTTC)                       |                                                                           |
| 0 Nov 2010    | Dr Notalia Paulis (NUUS)     | TD diagnostics discordant results                                         |
| 9 Nov.2018    | Dr Natalie Beylis (NHLS)     | TB diagnostics, discordant results, Heteroresistance and mixed infections |
|               |                              | (Natalie)                                                                 |
|               |                              | (Ivacane)                                                                 |

# **DTTC Academic Meetings 2018**

**The DTTC Academic Meetings 2018,** held fortnightly at the FMHS targeted academic researchers to engage in ground-breaking research initiatives. Speakers from local and abroad were invited to present their area of research in a broad range of relevant topics.

| Date         | Presenter          | Topic                                              |
|--------------|--------------------|----------------------------------------------------|
| 1 Feb. 2018  | Cornè Bosch        | Molecular diagnosis of M.TB from paediatric stool  |
|              |                    | samples – practice run for masters oral            |
| 8 Feb. 2018  | Dr. Theresa Mann   | Burden and trends of spinal tuberculosis in the    |
|              |                    | Western Cape                                       |
| 22 Feb. 2018 | Dr. Lena Ronge     | Open or closed Tuberculosis. "Are infants able to  |
|              |                    | spread TB?" - Findings from the European Childhood |
|              |                    | tuberculosis Archive                               |
| 1 Mar. 2018  | Prof. Helena Rabie | Paediatric HIV current challenges, future          |
|              |                    | opportunities                                      |
| 22 Mar. 2018 | Dr. Muhammad Osman | Paediatric XDR-TB Treatment outcomes – A           |

|              |                                                            | systematic review and individual patient data meta-<br>analysis.                                                                                                            |
|--------------|------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 05 Apr. 2018 | Dr. Florian Marx                                           | Former TB patients in the context of TB and HIV epidemics - need for tailored health interventions?                                                                         |
| 19 Apr. 2018 | Dr. Sue-Ann Meehan                                         | The contribution of a community based HIV counseling and testing (HCT) initiative in working towards increasing access to HIV counseling and testing in Cape Town           |
| 03 May 2018  | Dr. James Seddon                                           | The impact of household contact management on childhood tuberculosis: a mathematical modelling study                                                                        |
| 24 May 2018  | Dillon Wademan<br>&<br>Hanlie Myburgh                      | Chronic conditions and care: Intergenerational experiences of living with HIV, TB and Diabetes in South Africa (Dillon)                                                     |
|              |                                                            | Relationships, responsibility and the state: Understanding uptake of antiretroviral treatment in South Africa (Hanlie)                                                      |
| 31 May 2018  | Dr. Kyla Comins- TASK                                      | Update on MDR – TB treatment trials in adults                                                                                                                               |
| 28 Jun. 2018 | Dr. Florian Marx                                           | Projecting the population-levels effects of tuberculosis control in South Africa – update from the DTTC – WGDAM -Working group Data Analysis and Modeling                   |
| 05 Jul. 2018 | Dillon Wademan Lario Viljoen Laing de Villiers             | How health workers motivated People living with HIV (PLHIV) to initiate antiretroviral treatment (ART) at high CD4 counts within the HPTN 071 (PopART) trial (Dillon)       |
|              | Prof Kelly Dooley                                          | Public condition/private affliction: Negotiating comorbidity, confidentiality, and care in South Africa, a narrative from the HPTN 071 (PopART) study (Lario)               |
|              |                                                            | Stigma, secrecy, social support and HIV service access among transgender and gender-fluid people in South Africa, the HPTN 071 (PopART) study (Laing)                       |
|              |                                                            | Update on adult trials of TB/HIV (Kelly)                                                                                                                                    |
| 12 July      | Dr. Celeste De Vaal                                        | Raising ethical issues in the dead: An exploration of ethical challenges in Forensic Medicine                                                                               |
| 02 Aug. 2018 | Graeme Hoddinott and<br>Constance Mubekapi-<br>Musadaidzwa | HIV services for young people aged 10-24; priorities, gaps, and opportunities from the PopART for young people (P-ART-Y) study                                              |
| 16 August    | Dr Tony Garcia-Prats                                       | Results of the Bedaquiline CRUSH study     Emerging data on linezolid PK and safety in children with MDR-TB- feedback- Adult formulation tested ,tablets dissolve in water. |
| 23 Aug. 2018 | Prof Andrew Whitelaw                                       | S.areus epidemiology resistance virulance and diagnosis                                                                                                                     |

| 20 Sept. 2018 | Sociobehavioural        | 1)Acceptability of MDR-TB treatment for children,      |
|---------------|-------------------------|--------------------------------------------------------|
|               | science team            | findings from the MDR PK2 Masterchef.                  |
|               | presenters: Kyla        | 2)Video presentation of early thoughts on a            |
|               | Meyerson, Leletu        | knowledge, adherence, and social support app for       |
|               | Busakwe, and Stephanie  | children with TB.                                      |
|               | Jacobs.                 |                                                        |
| 04 Oct. 2018  | Dr. Amy Slogrove        | Evaluating long term child health outcomes following   |
|               |                         | pregnancy exposures in South Africa - using HIV and    |
|               |                         | antiretroviral drugs as a starting point"              |
| 18 Oct. 2018  | Dr. Elisa Lopez         | Pharmacokinetics of antituberculosis drugs at the site |
|               |                         | of disease in children with complicated intrathoracic  |
|               |                         | tuberculosis                                           |
| 08 Nov. 2018  | Dr Marije van Schalkwyk | CTU Academic Meeting: Drug PK in pregnancy, what       |
|               |                         | we know we don't know                                  |
| 29 Nov.2018   | Dr. Sue Purchase        | Looking through a different lens: TB in Literature and |
|               |                         | the Arts                                               |

# ADVOCACY AND COMMUNITY ENGAGEMENT

World TB Day is an annual joint initiative between the Stop TB Partnership and the WHO Global TB Programme to raise public awareness about the devastating impact of TB. On Friday 23<sup>rd</sup> March, DTTC organized a World TB Day lunch time event at the FMHS campus in collaboration with TB Proof and TB Alliance. The program included two interactive presentations: *TB prevention in children in the era of TB elimination* (Prof Anneke Hesseling) and *TB prevention in healthcare workers: Caring for the Carers* (Dr Angela Dramowski). The event concluded with a live interview with an adolescent MDR-TB research participant and her mother, explaining the MDR-TB treatment journey from a family perspective. Academic posters were displayed in the hallway for the duration of the day. DTTC staff and CAB members handed out TB information leaflets to FMHS staff and students during the morning to raise general awareness of TB on campus.







# The DTTC Community Advisory Board (DTTC CAB)

The establishment of a DTTC-CAB with a TB focus was initiated in June 2015 and was officially launched in November 2015, when the first general meeting was held at the Brooklyn Chest Hospital site. Since then, monthly General Meetings are conducted throughout the year. From 2016, the seven Steering Committee members held additional monthly meetings. DTTC CAB activities for January to December 2018 included:

**Meetings:** The DTTC CAB participated in General meetings; Steering Committee (SC) meetings for the executive body and SUN-CTU CAB Leadership Group (SCCLG) Meetings (i.e with DTTC, FAMCRU and TASK) and a representative participated in every IMPAACT ICAB bi-monthly teleconference call and HPTN CWG monthly teleconference calls including the HPTN 084 study monthly conference calls.

**CAB Annual Recruitment and Retention Drive**: 14 April 2018 the DTTC CAB hosted and "Open Day" and 22 prospective CAB members attended. Interviews were conducted during May and July. Subsequently, 4 adults and 10 youth were recruited into the DTTC CAB.

**HPTN 084 Protocol Review and Study Updates**: In February the CAB reviewed the protocol; and educational and marketing materials for the HPTN 084 study. The CLO met with the 084 Monitor to discuss CAB operations. The CAB and CLO met with 084 study staff to provide input; and were involved in activities relating to the call to pause the study for activation and enrolments and the activation.

**IMPAACT & HPTN Protocol Review and Study Updates:** On 09 June the Study Coordinator presented updates on P1101 and P1026. 14 June the CLO presented a CAB overview to the HPTN 084 team at the Kuilsriver site office.

**Network Training:** 04–05 February the CAB attended a training workshop hosted by FHI, "Understanding the Clinical Research Process". Network webinar training workshop invitations for various topics were disseminated and attended where possible.

**CAB Youth Development:** To raise the voice of our youth 2 youth members were elected to serve on the DTTC CAB Steering Committee during the 25 August Meeting. The youth group have written, produced and directed a 15 minute skit on HIV and TB incorporating a poem authored by one of the members and a song to illustrate the life of young people affected and infected with HIV and TB in the South African context. The production was presented at the 12<sup>th</sup> International Child TB Training Course. A 14 year-old youth member of the CAB who survived MDR-TB and her mom also shared their experiences with living with MDR-TB at the 12<sup>th</sup> International Child TB Training course and interacted with course participants thereafter.

International Network Meetings: 14 – 18 May the CAB Chairperson attended the HPTN Annual Meeting in Washington. No funds were available for community representation at the IMPAACT Annual Meeting in June 2018. A video production, "Two Countries; Two Choices" on TB by *Aids-Free World*, which featured 2 of DTTC adolescent research participants, one a CAB member, was screened at the International Aids Conference in Amsterdam, 23 – 27 July 2019.

The Annual Capacity Building Event & Award Ceremony: The CTU CABs (i.e. DTTC, FAMCRU and TASK) held their Annual Capacity Building Retreat on 18 -19 October 2018 at the Tygerberg Campus. This event culminated in the CAB Gala Dinner & Award Ceremony on Saturday, 20 October 2018 at the Bellville Civic Centre. This year we established an award with a floating trophy in honour of, and called the Katleho Mosimanegape Community Achievement Award, which recognizes outstanding leadership, commitment and dedication within the community forum. This year it was awarded posthumously to Katleho Mosimanegape, himself a TB and HIV survivor, who died in an accident on 27 June 2018. The Award will be held in DTTC for 12 months. Representatives of his family received the award at the Gala Dinner on 20 October 2019 at the Bellville Civic Centre.

**Annual Dissemination Meeting:** On 23 November 2018, the CAB members attended the Annual DTTC Dissemination Meeting, held at the Lord Charles Hotel Somerset West.

**DAIDS Programme Officer Site Visit**: On 04 September 2018, Eileen Pouliot, the DAIDS Programme Officer met with the community members during a joint CTU CAB Overview presentation at Tygerberg, J8 seminar room.

**World TB Day:** 23 March 2018, the CAB members participated in the Word TB Day programme hosted by DTTC at the Tygerberg campus.

**Special extramural events:** On 9 October 2018 12 members of the CAB attended the 8<sup>th</sup> International Desmond Tutu Peace Lecture at the Artscape where guest speaker, Cyril Ramaphosa, addressed the audience. On 20 October 2018 the CLO presented a CAB Overview to to US Health Attaché, Steven Smith at the Kuilsriver site office.

**The DTTC CAB Budget**: In March the CAB SC reviewed the 2018 CAB Budget which was accepted in principle by the DTTC EXCO in April.

**The 2019 IMPAACT Budget**: The CLOs of DTTC, FAMCRU and TASK met in November 2018 to finalise the 2019 IMPAACT Budget. The final proposal for the R178 000.00 allocation was submitted to the CTU Unit Manager and SU Grants Office on 7 December 2018.

**Funding Opportunity - Public engagement Fund**: The CAB CLO and two DTTC executive officers met to discuss the funding opportunity with the Wellcome Trust regarding a possible funding opportunity through creative approaches in engaging the public with health research.

**Condolences:** 27 June 2018, Executive member and Acting-Secretary, Katleho Mosimanegape, passed away in an accident. A vigil was held in the DTTC seminar room on 28 June 2018 and CAB members and DTTC staff visited the family home that evening. CTU CAB members attended the Memorial Service on 3 July 2018. The DTTC CAB and staff participated in a collection of cash and kind for the bereaved family as part of the 18 July 2018 Madiba Day activities.

#### The HPTN 071 (PopART) CAB

In the HPTN 071 (PopART) trial, each participating clinic had a health Committee that links it with its community. The PopART CAB was established in 2013 with representatives from the 9 sites in which PopART is being conducted. These representatives (2 from each site) were chosen from health committee members and they work as volunteers in making sure that health related issues within their communities and the clinics are addressed properly. The CAB is the link between the communities and the PopART researchers and has a constitution with rules and guidelines as to how to conduct their meetings. The PopART CAB meets once a month where various study related issues are discussed.

The CAB helps the researchers in making sure that community related issues are addressed and that misunderstandings between the community and the researchers are dealt with in a professional way. The CAB takes initiative in setting up these meetings. The CAB also helps in reviewing community related study material like informed consent documents and study questionnaires. In return, DTTC offers trainings for these members to develop them. The PopART CAB members have received training in the following areas:

- GCP training
- Basic HIV knowledge
- Minute and record keeping
- Basic Counselling
- Cancer screening tips

The PopART CAB, Community Engagement team and the Intervention Managers had a Workshop at the Lord Charles Hotel in preparation for the six events of Disseminating Intervention results planned for June /July 2018. The CAB and the teams worked together to find appropriate ways to communicate study results communities and making sure that closed venues are secured for the six dissemination events. The mobilisers are working closely with the CAB members in making sure that all stakeholders are aware of and invited to these events.

- The CAB members have been taking the lead in the establishment of community committees in preparation for disseminations (Intervention and the upcoming Primary Outcomes Dissemination Meeting).
- All the Nine Primary Outcomes Dissemination meetings will be held in March 2019.
- The role of CAB is to use community platforms to invite the relevant stakeholders
- The CAB meetings are held bi-monthly to review progress about these preparations.
- The last CAB for the year was on the 5<sup>th</sup> of November 2018 where site reports from all community committees were laid down to ensure the readiness of all sites for the Dissemination meeting.

# **HPTN 071 (PopART) Community Engagement (CE)**

The HPTN 071 (PopART) community engagement team were involved in several activities for the final year of the trial. These include the following:

In 2018, the Intervention District Logistic Managers (DLOs) and the Data Coordinators together with the Community Advisory Board members (CABs) for each site formed community working groups in preparation for the Intervention Dissemination meetings for the six communities involved.

- Out of the six, four dissemination meetings were held in June and the remaining two in July, 2018.
- At the end of the Population Cohort (PC) Research Assistant's contracts in June 2018, the DLOs and the CAB helped by members of the CE team started preparing for the Primary Outcomes Community Dissemination meetings to be held in March, 2019.
- Community Committees comprising of 2 CAB members, 1 CE member and 1 PC member were established to work on all site logistics for each site.
- The logistics included securing of venues, identifying community stakeholders to be invited and determining the most appropriate means to invite community members in the relevant sites
- The CE team was also consulting with the events office within the City of Cape Town to ensure that all events followed guidelines.

# The HPTN 071 (PopART) P-ART-Y Adolescent Community Advisory Board (ACAB)

The Adolescent CAB advised researchers on how best to implement the P-ART-Y evaluation survey and supported the community-level delivery of optimisations of the PopART intervention package for young people. In 2018 the P-ART- Y CAB members were involved in the general and focused meetings in which study processes and CAB issues were discussed.

The PARTY study team and five ACAB members (Luthando Ngwatyu, Nandipha Anta, Jafta Siyavuaya Nonkondlo, Katleho Mosimanengape and Mihlali Ngqawule) attend the Annual DTTC Dissemination Meeting held at Stellenbosch University on the 8 December 2017. This meeting brought together various stakeholders from the Western Cape Department of Health, City of Cape Town, and DTTC Paediatrics and Community Advisory boards. The meeting was themed "With Communities, for Communities", with aim of discussing studies currently underway at the centre and sharing lessons learnt. The ACAB had its first meeting on the 3rd of February 2018, and the meeting commenced with young people giving feedback on the DTTC annual dissemination meeting. The ACAB members then reviewed the dissemination plan and suggested ways of reaching young people. ACAB closeout was discussed and it was recommended that members apply to be part of the DTTC CAB. Since then, 7 P-ART-Y ACAB members have been welcomed into the DTTC CAB and continue to engage in CAB meetings.

#### **CONFERENCES**

During 2018, personnel from the DTTC participated in several local and international conferences, including the Union conference in The Hague, the International AIDS conference in Amsterdam, the South African TB conference in Durban, the Stellenbosch University research day and several other academic platforms.

# PEPFAR Conference. Pretoria, South Africa. 16 January 2018.

- HPTN 071 (PopART) staff presented study findings and lessons learnt

# SACEMA seminar. Stellenbosch, South Africa. 9 March 2018

- Sue-Ann Meehan. Community based HIV testing services: using a case study from Cape Town to provide evidence-based lessons learnt for program implementation.

# Annual TB Seminar at the Human Sciences Research Council. Cape Town, South Africa, 28 March 2018

- Boffa J. Is prevention better than cure? Perceptions of isoniazid preventive therapy in KwaZulu-Natal.

# <u>Peadiatric TB Pre-conference Symposium and Round Table Discussion at the 5<sup>th</sup> South African TB conference.</u> Durban. 12 June 2018

Convened and chaired by the DTTC: Tuberculosis in children: old problems and new solutions

- James Seddon. The evolving epidemiology of TB in children
- Karen du Preez. Children with TB in South Africa: where are they, who are they, and where do we find more of them.
- Marieke van der Zalm. Paediatric TB within the broader lung health context
- Anneke Hesseling. Therapeutic strategies for drug-susceptible and drug-resistant TB in children

# 5<sup>th</sup> South African TB Conference. Durban, South Africa. 12 - 15 June 2018

- Sue Purchase. Acceptability and palatability of a novel paediatric dispersible levofloxacin formulation in children

# Oral Presentations:

- Karen du Preez, Simon Schaaf, Rory Dunbar, Elisabetta Walters, Alvera Swartz, Regan Solomons, Anneke Hesseling. Complementary hospital-based surveillance strategies to better characterise the epidemiology of tuberculosis in children.
- Boffa J, Mayan M, Mhlaba T, Ndlovu S, Williamson T, Fisher D. Why agency is important when implementing IPT: Lessons from oMakoti in KwaZulu-Natal, South Africa.
- Boffa J, Mayan M, Ndlovu S, Fisher D, Staples S, Sauve R, Williamson T. Is prevention better than cure? Perceptions of isoniazid preventive therapy in KwaZulu-Natal, South Africa.
- Boffa J, Fisher D, Sauve R, Mayan M, Williamson T. The effectiveness of untargeted 6-month IPT to reduce tuberculosis incidence among people living with HIV with and without antiretroviral therapy in KwaZulu-Natal, South Africa.

# Poster presentation

- Boffa J, Mayan M, MacDonald H, Cele N, Dhladhla C, Ndlovu S. Creating culturally-appropriate dissemination products in tuberculosis research.

# AIDS 2018 Pre-conference and 4<sup>th</sup> International Conference on the Social Sciences and Humanities in HIV (ASSHH). Amsterdam, Netherlands. 20 – 23 July 2018.

# **Oral presentations**

L. Viljoen, L. Reynolds, H. Myburgh, G. Hoddinott, V. Bond, on behalf of the HPTN 071
 (PopART) team. Public condition/private affliction: negotiating co-morbidity, confidentiality,

- and care in South Africa, a case-study from the HPTN 071 (PopART) study.
- N. Peton, C. Mubekapi-Musadaidzwa, D. Wademan, P. Hendricks, G. Carolus, R. Mbaezue, J. Kruger, K. Jennings, C. Gunst, N. Grobbelaar, F. Louis, P. Bock, G. Hoddinott. How health workers motivated People living with HIV (PLHIV) to initiate antiretroviral treatment (ART) at high CD4 counts, the HPTN 071 (PopART) study.
- L de Villiers. L. Reynolds, A. Thomas, L. Busakwe, J. Hargreaves, A. Stangl, V. Bond, P. Bock, G. Hoddinott, on behalf of the HPTN 071 (PopART) study team. Stigma, secrecy, social support and HIV service access among transgender and gender-uid people in South Africa, the HPTN 071 (PopART) study.
- V. Bond, M. Mwamba, N. Vanqa, M. Steinhaus, D. Ziba, D. Milimo, L. Viljoen, T. Mainga, T. Nicholson, H. Ayles, S. Fidler, R. Hayes, P. Bock, J. Hargreaves, G. Hoddinott, J. Seeley and A. Stangl on behalf of the HPTN 071 (PopART) study team. "Self-stigma is the worst kind of stigma": Internalised stigma and navigating HIV care amongst health workers living with HIV in Zambia and South Africa, the HPTN 071 (PopART) stigma ancillary study.

# <u>22<sup>nd</sup> International AIDS Conference. Amsterdam, Netherlands. 23 – 27 July 2018.</u> *Oral presentations*

- G. Hoddinott on behalf of the HPTN 071 (PopART) study team. Acceptability of a HIV prevention package among young people: HPTN 071 (PopART) for Youth Study.

# Poster presentations

- S.-A. Meehan, D. Wademan, G. Hoddinott, R. Holtman, N. Grobbelaar, J. Kruger, M. Leonard, B. Yang, V. Naidoo, P. Bock, C. Gunst. Implementing community-based adherence clubs for stable HIV-infected patients in South Africa. Lessons learned from patient and health worker experiences.
- S.-A. Meehan, D. Wademan, G. Hoddinott, R. Holtman, N. Grobbelaar, J. Kruger, L. Maschilla, B. Yang, V. Naidoo, P. Bock, C. Gunst. Evaluating the impact of community-based adherence clubs on the clinical outcomes amongst ART patients in the Cape Winelands district of Western Cape Province of South Africa.
- S. Fidler, S. Floyd, B. Yang, M. Phiri, D. McLeod, A. Schaap, R. Sloot, K. Shanaube, S. Griffith, P. Bock, N. Beyers, H. Ayles, R. Hayes, on behalf of the HPTN071(PopART) Study Team.
   Dramatic reductions in time to ART initiation among HIV+ individuals referred to HIV care following home-based testing services: Experiences from the HPTN071(PopART) trial between 2014 and 2017.
- K. Shanaube, D. Macleod, J. Mwate Chaila, G. Hoddinott, A. Schaap, S. Floyd, C. Mackworth-Young, P. Bock, R. Hayes, S. Fidler, H. Ayles, HPTN071 (PopART) Study Team. HIV care cascade among adolescents in a 'test and treat' community based intervention in Zambia and South Africa: HPTN071 (PopART) for Youth study.
- R. Thomas, S. Kanema, L. Mwenge, S. Floyd, P. Bock, H. Ayles, N. Beyers, S. Fidler, R. Hayes,
   K. Hauck, HPTN 071(PopART) Study Team. Costs of two rounds of home-based HIV testing
   and counselling in Zambia: Evidence from the HPTN 071(PopART) study.
- R. Thomas, R. Friebel, K. Barker, L. Mwenge, S. Kanema, N. Vanqa, A. Harper, N. Bell-Mandla, P. Smith, S. Floyd, P. Bock, H. Ayles, N. Beyers, S. Fidler, R. Hayes, K. Hauck, HPTN 071(PopART) Study Team. Work and home productivity of HIV-positive and HIV-negative individuals in Zambia and South Africa: A cross-sectional baseline survey of the HPTN071/PopART trial.
- M. Simwinga, J. Mwate, T. Ng'ombe, S. Belemu, N. Makola, C. Mubekapi-Musadaidzwa, G. Hoddinott, R. White, K. Shanaube, V. Bond, T. Mainga. Adolescent and young people's

- participation and representation in clinical trials: Lessons from a community-wide HIV testing and treatment study, the HPTN 071 (PopART) study. (poster discussion).
- M. Mbewe, C. Mackworth-Young, D. Ziba, R. Ndubani1, B. Chiti, M. Simuyaba, M. Gondwe, K. Shanaube, G. Hoddinott, V. Bond. 'In their shoes': Limited disclosure amongst young people living with HIV stifles their ability to contact others in the same situation, a qualitative cohort HPTN071 (PopART) study in Zambia.
- D. Macleod, B. Yang, S. Floyd, A. Schaap, S. Griffith4, K. Shanaube, M. Phiri, R. Sloot, P. Bock, N. Beyers, H. Ayles, S. Fidler, R. Hayes. Retention on ART within the HPTN 071 (PopART) universal testing and treatment programme in Zambia and South Africa.
- J. Hargreaves, A. Stangl, G. Hoddinott, T. Mainga, S. Floyd, K. Sabapathy, H. Jones, L. Viljoen, T. Pliakas, S. Krishnaratne, C. Mubekapi-Musadaidzwa, V. Bond, HPTN (071) PopART Team. The HIV stigma landscape in sub-Saharan Africa: Baseline findings of a mixed-method, comprehensive evaluation nested in the HPTN 071 (PopART) trial.
- R.J. Hayes, S. Floyd, A. Schaap, K. Shanaube, B. Yang, S. Griffith, P. Bock, N. Beyers, H. Ayles, S. Fidler, HPTN 071 (PopART) Study Team. Achieving the first two UNAIDS 90-90-90 targets on completion of a three-year universal testing and treatment (UTT) intervention in the HPTN 071 (PopART) randomised trial in Zambia and South Africa.
- B. Yang, R. Sloot, S. Floyd, F. Esau1, J. Molaolwa1, D. Awoniyi, Y. Saunders1, P. Siboto, L. Gqiba-Tunywa, D. Myburgh, B. Cimi, S. Griffith, P. Bock, H. Ayles, S. Fidler, R. Hayes, N. Beyers, HPTN 071 (PopART) Study Team. HIV testing patterns in two community-based approaches to universal test and treat in the HPTN 071 (PopART) intervention in South Africa.
- K. Sabapathy, C. Mulubwa, C. Mubekapi-Musadaidzwa, V. Bond, G. Hoddinott, H. Stockl, S. Floyd, P. Bock, J. Seeley, H. Ayles, S. Fidler, R. Hayes, HPTN 071 (PopART) Study Group.
   Intimate partner violence and associated factors in HPTN 071 (PopART) study communities a comparison by HIV status.

# <u>Stellenbosch University 6nd Annual Academic Day. Tygerberg Medical Campus. 29 August 2018.</u> *Oral presentations*

- Celeste de Waal. Raising ethical issues in the dead: An exploration of ethical challenges in forensic medicine
- Peter Bock on behalf of the HPTN 071 (PopART) team. Baseline CD 4 count as a predictor of clinical outcomes when providing antiretroviral treatment regardless of CD 4 count at government clinics within the HPTN 071 (PopART) trial.
- Lario Viljoen on behalf of the HPTN 071 (PopART) team. Public condition/ private affliction: Negotiating comorbidity, confidentiality and care in South Africa, a case study from the HPTN 071 (PopART) study.
- Karen du Preez. Ten years of routine paediatric TB surveillance data in South Africa high disease burden and poor treatment outcomes

#### Poster presentations

- Sue-Ann Meehan. Evaluating the impact of adherence clubs on the clinical outcome amongst ART patients in the Cape Winelands District of Western Cape Province, South Africa
- Dillon Wademan. Implementing community-based adherence clubs for stable HIV-Infected patients in South Africa. Lessons learned from patient sand health worker experiences.
- Blia Yang. HIV-testing patterns in two community-based approaches to Universal Test and Treat in the HPTN 071 (PopART) intervention in South Africa.

- Mareli Claassens. Mortality amongst HIV-positive female tuberculosis patients in South Africa.
- Constance Mubekapi-Musadaidzwa. Motivating people living with HIV to initiate antiretroviral treatment at high CD 4 counts in three HPTN 071 (PopART) health facilities in the Western Cape, South Africa.
- Kerry Nel. Patterns of facility-based HIV testing service (HTS) at two public health facilities participating in the HPTN 071 (PopART) Universal Test and Treat intervention in Cape Town, South Africa.
- Mark Theart. Using aerial photography in longitudinal community studies
- Jana Winckler. The pharmacokinetics of high dose isoniazid for the prevention or treatment of drug-resistant tuberculosis in HIV-infected and –uninfected children (The SHINE trial)
- Megan Palmer. Use of chest radiographs in a phase 3 randomised controlled efficacy trial for treatment shortening of non-severe drug-susceptible TB in children (The SHINE trial)
- Klassina Zimri. Clinical trial and emotional labour: The role of the research nurse in paediatric MDR-TB studies
- Louwina van der Laan. Pharmacokinetics of intracellular Stavudine-Triphosphate in children after reduced dose: Can we improve Stavudine's safety profile?
- Nomsa Apleni, Gerald J Maarman, Nomtha Mandla, Peter Bock, Nulda Beyers on behalf of the HPTN 071 (PopART) team. Presenter: Kate Fortuin. Roles of nurses in ensuring quality and effectiveness of blood sample collection.

# SACEMA Research Conference. Stellenbosch. 10-13 September 2018

#### Poster presentation

- Muhammad Osman, Alex Welte, Rosemary Brown, Rory Dunbar, Graeme Hoddinott, Anneke C Hesseling, Florian M Marx. Former tuberculosis patients, a high-risk group for tuberculosis-associated morbidity and mortality in Cape Town, South Africa: a pilot study.

# 12th Annual Early Career Scientist Convention (ECSC). 17-19 October 2018

# Abstract published

- Muhammad Osman, Alex Welte, Rosemary Brown, Rory Dunbar, Graeme Hoddinott, Anneke C Hesseling, Florian M Marx. Former tuberculosis patients, a high-risk group for tuberculosis-associated morbidity and mortality in Cape Town, South Africa: a pilot study.

# <u>The 49TH Union World Conference on Lung Health. The Hague, Netherlands.</u> 24-27 October 2018. <u>Symposium organizer and chair</u>

- M. van der Zalm. Paediatric lung function measurements in low-middle income countries with a high burden of TB/ HIV, malnutrition and environmental exposures.

# Symposium presentation

- N Suryavanshi, G. Hoddinott High-quality social science to inform the design and interpretation of paediatric tuberculosis treatment trials

#### **Oral presentations**

- M. van der Zalm. Speaker at symposium: "Lung function in children presenting with symptoms suggestive of pulmonary tuberculosis".
- Garcia-Prats. Pharmacokinetics and pharmacodynamics in designing clinical trials in children
- Garcia-Prats. The current landscape of paediatric trials for the prevention and treatment of DS- and DR-TB: feeding the pipeline

- Garcia-Prats. Meet the Expert Session: Injectable-free regimens, and the role of injectable agents, in the treatment of children with DR-TB
- Garcia-Prats. Pharmacokinetics, safety and optimal dosing of linezolid in children with multidrug-resistant tuberculosis
- Garcia-Prats. Pharmacokinetics, safety and dosing of levofloxacin 100 mg dispersible tablets in children exposed to multidrug-resistant tuberculosis
- S. Schaaff. TB prevention in children in high TB burden settings: implementing the child contact management (CCM) care cascade. Introductory & 2nd talk: Preventive treatment considerations for multidrug-resistant tuberculosis
- S. Schaaff. Update on clinical and programmatic management of multidrug- and extensively drug-resistant TB (MDR-/XDR-TB) oral treatment of MDR-TB in children
- M Osman, EP Harausz, A Garcia-Prats, HS Schaaf, JA Seddon, AC Hesseling. Good treatment outcomes in children with extensively drug-resistant tuberculosis: a systematic review and individual patient meta-analysis.
- Megan Palmer Use of existing tools for entry points and end points in paediatric TB trials: how can we better use the chest radiograph?
- Elisabetta Walters, Pamela Nabeta, Padmapriya Banada, Anne-Marie Demers, Corne Bosch, Gary Reubenson, Naazley Pandor and Lesley Scott. Mycobacterium tuberculosis detection from stool in young children using a novel centrifugation-free method with Xpert MTB/RIF.

#### Short oral abstract

- M. Osman. Mortality amongst HIV-positive female tuberculosis patients in South Africa
- K. Du Preez M. Osman, P. Naidoo, S. Dlamini, A.C. Hesseling. Ten years of routine paediatric TB surveillance data in South Africa high disease burden and poor treatment outcomes
- F. Solomon-da Costa, K. Joseph, M. Osman, K. du Preez. Burden, characteristics and treatment outcomes of patients treated for Isoniazid mono-resistant TB in Cape Town, South Africa.

#### Poster presentations

- S. Schaaf. Culture-confirmed tuberculosis in infants less than 3 months of age: clinical presentation and management
- M Osman, K du Preez, R Dunbar, A Welte, P Naidoo, AC Hesseling. Trends in the South African tuberculosis epidemic with scale-up of TB-HIV integration services.
- M Palmer, E Walters, M vd Zalm, HR Draper, HS Schaaf, P Goussard, J Morrison, RP Gie, AC Hesseling. Could 'non-typical' chest radiographic features in children with culture-confirmed tuberculosis reflect early, less severe disease?
- M Palmer, R Mboizi, V Mulenga, A Kinikar, D Baskaran, M Thomason, A Crook, AC Hesseling and the SHINE trial team. Use of chest radiographs in a phase 3 randomised controlled efficacy trial for treatment shortening of non-severe drug-susceptible TB in children (the SHINE Trial)

# SCIENTIFIC MEETINGS, WORKSHOPS AND EVENTS

# January 2018

- GATES meeting: Demo project to address the gaps in TB testing
- National TB Network SA Focus on BRIX Countries for TB trials (Anneke Hesseling)

#### February 2018

- Academic Writing Workshop (Blia Yang, Nomtha Mandla, Kerry Nel, Constance Mubekapi-Musdaidzwa, Rosa Sloot, Mohammed Osman, Gerald Maarman)
- FHI 360 Workshop: Understanding the Clinical Research Process and Principles of Clinical Research (PopART team)
- HPTN071 (PopART) annual workshop in Cape Town. (PopART team)
- Gates Foundation ILTF meeting
- IMPAACT 2005 Regional Meeting

#### March 2018

- IMPAACT Smart-kids P2020 protocol writing workshop (Anneke Hesseling, Simon Schaaf, James Seddon, Tony Garcia-Prats)
- PhD Graduation gala event (Peter Bock, Florian Marx, Sue-Ann Meehan and Rory Dunbar)
- CAB General Meeting: planning of open day to recruit additional members (Gwynneth Hendricks; DTTC CAB & prospective members)
- South African Social Science and HIV Programme (SASH) event: What does the future of HIV research look like? (Sociobehavioural science team)
- Department of Health in Winelands workshop: lessons learnt from field staff for transmission into policy - budget concerns (PopART team)
- WHO Meeting in Geneva (Anne-Marie Demers, Anneke Hesseling)
- Stigma meeting: Dissemination meeting on clinical level. Fieldworker (CHiPs) data feedback to healthcare services (Kerry Nel)
- World TB Day Event on Tygerberg Campus (DTTC staff and Tygerberg campus staff)
- Gates Foundation ILTFU meeting in Johannesburg (Anneke Hesseling, Muhammad Osman)
- Paediatric TB Meeting in Lithuania (James Seddon, Tony Garcia-Prats)
- Final P-ART-Y CAB meeting before members integrated into DTTC CAB

#### **April 2018**

- Mini ORAP Workshop (facilitated by Muhammad Osman)
- Thoracic Meeting in Johannesburg presentations on preventative therapy and diagnosis in children (Anneke Hesseling, Muhammad Osman)
- CAB Meeting: Recrutiment and Retention Drive
- Various research visits: Florian Marx to Europe; Anne-Marie Demers to Peru; James Seddon to the United Kingdom
- APSA meeting in Johannesburg (Constance Mubekapi-Musdadzwa, Graeme Hoddinott)
- HPTN 071 (PopART) Secondary outcomes meeting with Zambian Colleagues (Sue-Ann Meehan)
- HPTN 084 Lab Audit

### May 2018

- AURUM Global Funding Meeting in Johannesburg (Anneke Hesseling)
- HPTN Annual Meeting in Washington, USA. (Sue-Ann Meehan, Peter Bock, Blia Yang, Nomtha Mandla, Redwaan Vermaak, Nozi Makola, Jabulile Mantantana, and 2 CAB Members)

- HPTN Annual Community Working Group Meeting in Washington, USA (Nozi Makola)
- Scientific Advisory Board Meeting in Paris. Workshop: Chest X-Rays in Children (James Seddon)
- Research Day plenary presentation (Nulda Beyers)
- Pre-conference Union Planning Meeting in the Hague (James Seddon)
- TB Think Tank Meeting (Johannesburg). LTFU finding the missing cases. Improvement of health info exchange by upgrading the HIE for easier data reporting (Muhammad Osman)
- Research Day at UKWANDA Stellenbosch University Worcester Campus. Nulda Beyers presented.
- Scientific Advisory Board Meeting in Paris: 1st TB-Speed symposium (re TB in children)

#### **June 2018**

- IMPAACT Annual Meeting in Washington. (Tony Garcia-Prats, Frieda Verheye-Dua, Anne-Marie Demers, Sharon Mbaba, Simon Schaaf, Anneke Hesseling, Anneen van Deventer)
- Community Dissemination of HPTN071 (PopART)
- CAB general meeting: HPTN meeting feedback and study protocol reviews.
- British academy writing retreat at Goedgedacht farm (Malmesbury). Six of the Sociobehavioural Science Team received sponsorships: Graeme Hoddinott, Laing de Villiers, Dionne Jivan, Rozanne Casper, Nosivuyile Vanqa and Angelique Thomas.
- SUN-CTU Meeting chaired by Prof Mark Cotton.
- ACTG Network Meeting in Washington DC (Annemarie Demers).
- Consultation workshop for Scoping review on ART scale-up in South Africa with the PopART sociobehavioural science team in collaboration with HIV programme implementers and managers at provincial, city, and district levels (Hanlie Myburgh).
- GHIT Funding Presentation Tokyo, Japan (Anneke Hesseling).
- Scalabrini The team addressed and shared information with foreigners on TB and HIV, chronic illnesses, access to clinics and services available if unemployed.

# July 2018

- KNCV MDR-TB Training Roll Out of preventative therapy for MDR-TB exposed children Indonesia (Presenter: Tony Garcia-Prats).
- HPTN 071 (PopART) Stigma working group workshop at London School of Hygiene and Tropical Medicine (Graeme Hoddinott, Lario Viljoen, Constance Mubekapi-Musadaidzwa, Laing de Villiers, Dillon Wademan).

# August 2018

- Cluster Randomized Trials for Evaluation of HIV interventions workshop at FMHS. Facilitated by Prof. Rhoderick Machekano.
- Marieke van der Zalm visited the Wilhelmina Children's' Hospital from 22 August to 30 October 2018 for capacity building in Lung Health as part of the HB Thom award received at the end of 2017.
- Graeme Hoddinott and Karen du Preez attended and a multi-stakeholder meeting on behalf of DTTC hosted by TB HIV Care for the South African Assessment of Barriers & Facilitators to TB Services in Johannesburg.

# September 2018

- 12<sup>th</sup> International Child TB Training Course at Goudini ATKV Spa, Rawsonville, Cape.
- SHINE TGM Meeting in Lusaka, Zambia (DTTC Paedicatric team)
- Faith Leaders meeting at the Kwazulu Natal Christian Council in Pietermaritzburg (Jody Boffa).
- Basic HIV counselling and testing short course (SETA). US hosted at Ruslamere in Durbanville.
   Councellors from DTTC Sociobehavioural science and Peadiatric teams CPUT and FAMCRU attended.
- Current TB meeting Adult TB. Complications after treatment and cure. (James Seddon).
- CHEC Module 1.8 Effective Communication. Devon Valley Hotel (Gwynneth Hendricks).
- TB meeting between Sociobehavioural science and Zambart Quantitative Data and Qualitative data score and Impact on study outcomes. (Graeme Hoddinott).

#### October 2018

- SACEMA meeting with Muhammad Osman presenting his paper (Muhammad Osman and Anneke Hesseling).
- Brittish Academic Writing workshop in Goedgedacht, Riebeeck Kasteel (Laing de Villiers; Dionne Jivan; Angelique Thomas, Rozanne Casper, Nosi Vanqa).
- Dr. Karen du Preez was selected to attend a World Health Organization Training Workshop for TB Surveillance: Establishing a Roster of Consultants Providing Technical Assistance for Epidemiological Reviews and Data Analysis and use in The African Region, which was held in Kigali, Rwanda (8 to 12 October 2018).
- CTU-CAB Retreat (FMHS). Two day training program involving TASK, FAMCRU and DTTC. A Gala Dinner was held on Saturday 20th Oct at the Bellville Civic Centre. In honour of Katleho the "Katleho Mosimanegape CAB Achievement Award" was established.

#### **November 2018**

- Discussions with WHO country representative on Operational Research, Surveillance and ETR.net .in Pretoria (Karen Du Preez)
- Paediatrics and Child Health Academic Day of Excellence (Liz Walters presented on Pulmonary TB)
- Desmond Tutu TB Centre Dissemination meeting at the Lord Charles Hotel. Opening guest Speaker is Professor Mark Cotton, FAMCRU, Department of Peadiatrics and Child Health and the topic of his talk, "The state of HIV & Prevention studies in South Africa.
- HPTN 071 (PopART) Stigma Working Group writing workshop in Choma, Zambia (Lario Viljoen, Dillon Wademan, Laing de Villiers, Angelique Thomas, Rozanne Casper)
- LINKEDin study site visits to Gauteng and KwaZulu Natal (Sue-Ann Meehan and Muhammad Osman). Visited NICD, spent time with implementing study partners (Right to Care and IRD), visiting hospitals and a selection of primary healthcare facilities in the Johannesburg Metro and the Ugu district, where the study will be implemented.
- WHO GIC Implementation Guidelines. Dr. James Seddon presented on Implementation of WHO guidelines for the treatment of MDR treatment in children.
- TB Innovations Consortium Meeting in Johannesburg (Anneke Hesseling; Sue-Ann Meehan; Muhammad Osman)

#### December 2018

- HPTN 071 (PopART) results "unblinding" meeting in Seattle, United States of Ameica (Peter

Bock)

# Ongoing 2018

- Joint SUN-CTU CAB Leadership Group Meetings (DTTC CAB EXCO; TASK CAB EXCO; FAMCRU EXCO). Meetings held throughout the year.
- Working Group for Data Analysis & Modeling (WGDAM). WGDAM is a working that aims to produce research output (such as our publication in SAMJ August 2018). They are currently working on a paper led by Muhammad Osman and Florian Marx, which aims to assess the impact of antiretroviral therapy on tuberculosis incidence and mortality in nine South African Provinces using a mathematical model. The group met on a regular basis throughout the year.

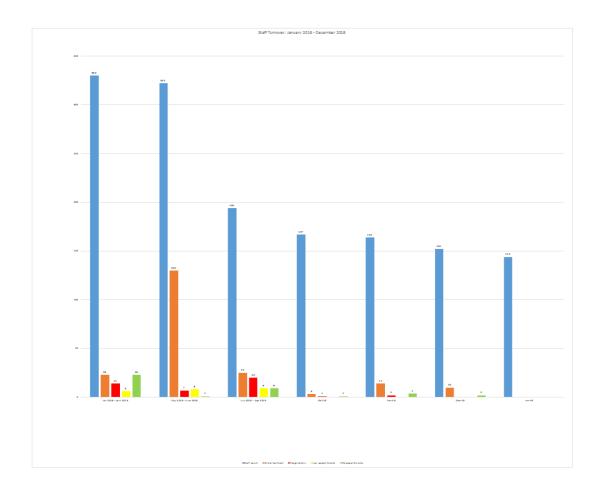
#### **VISITORS AT THE DTTC DURING 2018**

| Date            | Visitor                             | Reason                                  |
|-----------------|-------------------------------------|-----------------------------------------|
| 29 Jan. 2018    | Yael Hirsh-Moverman - Assistant     | Dr Yael Hirsch-Moverman visited the     |
|                 | Professor in Epidemiology (In ICAP) | paediatric team to discuss ongoing      |
|                 | at the Columbia University Medical  | collaboration and a joint NIH R01 grant |
|                 | Center, Mailman School of Public    | application that was submitted in June  |
|                 | Health, Columbia University.        | 2018.                                   |
| 5-9 Feb. 2018   | Dr Virginia Bond – Social Scientist | Discussion with Sociobehavioural        |
|                 | at ZAMBART in Zambia                | Science Team                            |
| 12 Feb. 2018    | Vundli Ramokolo – Senior Scientist  | Discussion with Anneke Hesseling        |
|                 | DOH Systems Research Unit, SA MRC   | regarding potential post-doc with DTTC  |
| 12-15 Feb. 2018 | Lily Telisinghe - London School of  | Visiting PopART, to work on TB case     |
|                 | Hygiene & Tropical Medicine         | notification data from PopART.          |
| 19-23 Feb. 2018 | Georgia White – AIDS Free World     | Discusison with Anneke regarding        |
|                 |                                     | filming in SA for TB/HIV advocacy       |
| Feb. 2018       | Carl Swenson – UPSALLA              | Discussion regarding TB dosing          |
|                 | Pharmacokinetics                    |                                         |
| Mar. 2018       | Dr Rafael Saulter and Dr William    | IMPAACT Smart-kids P2020 protocol       |
|                 | Probert, Researchers in Infectious  | writing workshop                        |
|                 | Disease Modelling Big Data          |                                         |
|                 | Institute, Ki La Shing Centre for   |                                         |
|                 | Health Information and Discovery,   |                                         |
|                 | Oxford University                   |                                         |
| Mar. 2018       | Pauline Howell – Senior Medical     | IMPAACT Smart-kids P2020 protocol       |
|                 | Officer at Sizwe Tropical Disease   | writing workshop                        |
|                 | Hospital                            |                                         |
|                 | Katie McCarthy - IMPAACT            |                                         |
|                 | Operations Centre, Science          |                                         |
|                 | Facilitation FHI360)                |                                         |
|                 | Leavitt Morrison, Soyeon Kim –      |                                         |
|                 | Harvard University                  |                                         |
| 16 Mar. 2018    | Jeff Haskins – Otsuka               | Meetings with Tony Garcia-Prats and     |
| 00000           |                                     | Anneke Hesseling                        |
| 26-29 Mar. 2018 | Yael Hirsch – Assistant Professor   | Meeting with Paediatric team            |
|                 | Epidemiology, Columbia University   |                                         |
| 0.40.4          | Medical Centre                      | 1                                       |
| 9-13 Apr.2018   | Ben Daniels – McGill University's   | Assisted Jody Bofta to set up a         |
|                 | Standardized Patients Project       | Quantitative Data Base                  |

| 17-18 Apr.2018   | Ms Mwelwa Phiri and Ms Ramya<br>Kumar                                                                                                              | HPTN 071 (PopART) visit                                                                                                   |
|------------------|----------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------|
| Jun Aug. 2018    | Rachel Morse – student at Kings<br>College London                                                                                                  | Exchange program through partnership between faculty and Department of Social Health and Medicine                         |
| 06-13 Jul. 2018  | Kelly Dooley - John Hopkins University and extraordinary professor at DTTC (Department of Paediatrics)                                             | Pharmacokinetic research meetings / Goudini TB Workshop                                                                   |
| 06-21 Jul. 2018  | Ava Archery (student)                                                                                                                              | Intern on SHINE study                                                                                                     |
| 06-13 Jul. 2018  | Sebastian Goodwin-Groen (student)                                                                                                                  | Intern                                                                                                                    |
| 03-07 Sept. 2018 | Oladapo Alli is PAB senior protocol pharmacist at the Division of AIDS.                                                                            | DAIDS site visit                                                                                                          |
| 03-07 Sept. 2018 | Eileen Pouliot - Branch Chief for<br>the Africa and Domestic Partners<br>Branch in the Office of Clinical Site<br>Oversight (OCSO)                 | DAIDS site visit / CTU visit                                                                                              |
| 17-19 Sept. 2018 | Erin Sizemore ; Rosanna Boyd   TB<br>Trials Consortium (TBTC)                                                                                      | Site Visit and training                                                                                                   |
| 28 Sept. 2018    | Teja Stojiljkovic, BARC                                                                                                                            | Site Visit                                                                                                                |
| 03-06 Sept. 2018 | Lebogang Tshehla                                                                                                                                   | Regulatory monitoring visit                                                                                               |
| 03-06 Sept. 2018 | PPD (IMPAACT)                                                                                                                                      | External monitoring visit                                                                                                 |
| 22-26 Oct. 2018  | Representatives of Consul General USA, Steven Smit (Health Attaché)                                                                                | Visiting Kuils River Site office                                                                                          |
| 30-31 Oct. 2018  | Cathy Tupman, Global Health Manager and Aimi Vanden-Oever Section Manager for Global Health   Imperial College, London- James Seddon               | Meetings with key academics and the<br>Grants office; DTTC, Tygerberg<br>Hospital, Brooklynn Chest Hospital               |
| 28-30 Nov. 2018  | PPD Monitor                                                                                                                                        | Monitoring visits at Kuils River and the DTTC                                                                             |
| 28–30 Nov. 2018  | Dessislava Tarlton (UNITAID) & Ekaterina Rykovanova (WHO)                                                                                          | Evaluation of the Centre and a walk through                                                                               |
| 31 Nov. 2018     | Representatives of Consul General USA, Steven Smit (Health Attaché) and Virginia Blaser  Visited the PopART team to assess project progress on the |                                                                                                                           |
| 28-30 Nov. 2018  | WHO-Unitaid (Led by Tony Garcia-<br>Prats)                                                                                                         | Visit to PK unit and Faculty doing an evaluation of facility, resources and staff. In depth assessment of all activities. |

# **EMPLOYEES**

During 2018 a total of (205) employee contracts ended. The exit of staff included the final annual round of the PC Field remunerators in June 2018. Between July and September, a high number of employees (20) resigned, seeking permanent employment elsewhere. The Centre also employed total of (23) new staff members on different studies, linked to different post levels. Total of (40) staff members could be re-employed, which relates our current HR retention strategy.



| Month           | Staff count | End of   | Resignations | New appoint. | Re-appoint. |
|-----------------|-------------|----------|--------------|--------------|-------------|
|                 |             | contract |              |              |             |
| Jan – Apr. 2018 | 330         | 23       | 14           | 6            | 23          |
| May - Jun. 2018 | 322         | 130      | 7            | 8            | 1           |
| Jul Sept. 2018  | 194         | 25       | 20           | 9            | 9           |
| Oct. 2018       | 167         | 3        | 1            | -            | 1           |
| Nov. 2018       | 164         | 14       | 2            | -            | 4           |
| Dec. 2018       | 152         | 10       | -            | -            | 2           |
| Jan. 2019       | 144         | -        | -            | -            | -           |

#### **DTTC GOVERNANCE**

# **DTTC Executive Committee (ExCo)**

The DTTC is led by the Director, Distinguished Professor Anneke Hesseling, who is assisted by an Executive Committee (EXCO) of 12 senior scientific and core support DTTC staff members. The Director, who chairs the EXCO, reports to the Executive Head of the Department of Paediatrics and Child Health, Faculty of Medicine and Health Sciences as well as the DTTC Governing Board, which is responsible for oversight of the DTTC's strategic management. The governing board is chaired by Professor Andrew Whitelaw, Dept. Medical Microbiology and NHLS, Tygerberg; vice-chair: Professor Wolfgang Preiser: Division Head: Medical Virology. All members of the ExCo committee play an active role in the management of DTTC, making key decisions to ensure the vision and mission of DTTC are maintained and expanded. Members also form part of the scientific strategy team and are responsible for driving the research agenda at DTTC forward, in line with the 3 DTTC research pillars. The role of this team is to provide scientific input into research ideas put forward to the team and collaborate on potential research proposals.

The DTTC ExCo meets once a month. In addition, there is a dedicated scientific strategy meeting convened once a month for the ExCo and additional *ad hoc* member to discuss and plan ongoing and future research priorities and strategies.

Primary focus areas for 2019 will include applying to remain within the DAIDS and TBTC networks which offer core and study specific funding over a 7 year period. In addition DTTC is actively applying for research opportunities to strengthen core funding to support studies being conducted across the DTTC platform.

#### DTTC EXECUTIVE COMMITTEE ROLES AND RESPONSIBILITIES

| Name                    | Role                                                          |
|-------------------------|---------------------------------------------------------------|
| Prof Anneke Hesseling   | DTTC Director and chair of DTTC ExCo                          |
| Dr Tony Garcia- Pratts  | Medical director of the Desmond Tutu TB Centre Brooklyn Chest |
|                         | Hospital Paediatric Pharmacokinetics Unit                     |
| Dr Marieke van der Zalm | Clinical lead: Paediatric lung health studies                 |
| Dr Frieda Verheye-Dua   | Regulatory Manager                                            |
| Prof Simon Schaaf       | Lead investigator Paediatric MDR-TB studies                   |
| Dr Sue-Ann Meehan       | Project Manager and co-investigator                           |
| Dr Peter Bock           | Lead HIV Prevention research                                  |
| Dr Muhammad Osman       | Lead: Implementation Research                                 |
| Mr Graeme Hoddinott     | Lead Sociobehavioural Scientist                               |
| Mrs Sterna Brand        | Human Resources Manager                                       |
| Mr Theo Smith           | Logistics /Operations Manager                                 |

| Dr Rory Dunbar      | Data centre manager |
|---------------------|---------------------|
| Dr AnneMarie Demers | Laboratory Director |

#### RISK MANAGEMENT

# Risks related to Human Resources, Finance and Regulatory are being monitored and evaluated on a regular basis.

The DTTC regulatory office will ensure that no study is commenced without appropriate regulatory approval.

The protection of study participant confidentiality remains a high priority for DTTC in our aim to deliver high quality reputable research outputs. DTTC will continue to support staff training in good clinical practice (GCP) and research confidentiality prior to any study commencement. Through constant data monitoring by dedicated data centre staff, duel entry of any research information as well as GCP training, DTTC will continue to mitigate against data fabrication. The regulatory office will also monitor the import of drugs or export of any materials to research partners to ensure no transfer is commenced without appropriate documentation.

Poor marketing and communication to external stakeholders, has resulted in low exposure to the scientific community and funding agencies. DTTC is actively working on a development plan to improve their marketing and communication efforts in order to improve our local and global footprint and develop strategic relationships with new and current external stakeholders. Accessibility to communities for recruitment and retention of study participants is still a potential risk. DTTC Field offices have been established in various communities and the development of community advisory boards to strengthen the support for DTTC in communities has been implemented to improve accessibility. Strong line management in field offices mitigates against any HR irregularities (attendance, working hours) that may arise.

Sustainability of core support staff is an ongoing risk due to a luck of core funding. Ongoing efforts to obtain core funding is being implemented. Human resource risks relating to the safety of staff training, staff contracts to retain core staff and staff diversity has been reduced.

Political instability and violence or unrest in study communities places staff and patient safety at risk. DTTC has implemented many safety measures but have also identified various other measures to provide safe working conditions for staff. In the first quarter of 2019 all DTTC drivers will be trained and retrained in defensive driving, including highjack prevention and evaluation. The high burden of TB disease and HIV infection among the population served, poses a health risk to DTTC staff. DTTC has taken many infection control measures through modification of vehicles and offices. Infection prevention control (IPC) training has been scheduled to start in February 2019, the aim is to train all DTTC staff in Infection prevention Control measurements. A Baseline Hep B testing campaign for all staff was launched in November 2018. Staff will be tested and vaccinated in line with national guidelines.

To mitigate the risks of temporary staff contracts leading to insecurity, demotivation and often to early exit, the DTTC has adapt all adverts too two year contracts with benefits, unless the appointment is for a specific intervention linked to a shorter term...

DTTC is committed to adhere to SU diversity policies during their recruitment processes. Staff has been identified from disadvantaged backgrounds to be promoted to levels that are more senior. Enrolment of Principle investigators in formal supervision programs has been implemented to support Master's and PhD students.

DTTC employs a cadre of finance staff who work closely with the faculty research grants management office (RGMO) to ensure no misappropriation of research funds takes place. Regular expenditure allocation, reporting and finance meetings are held to mitigate against any misappropriation. Correct budgeting in collaboration with RGMO will also mitigate against large exchange rate loss

#### ACKNOWLEDGEMENTS AND FUNDERS









































Health solutions for the poor

International Union Against

Tuberculosis and Lung Disease



TB ALLIANCE

GLOBAL ALLIANCE FOR TB DRUG DEVELOPMENT



























# South African Collaborators | Funders | Partners





CITY OF CAPE TOWN ISIXEKO SASEKAPA STAD KAAPSTAD





SARChI Chair in Paediatric TB (Hesseling)





Department for International Development















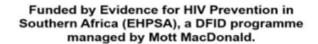
















#### **APPENDIX I. RESEARCH OUTPUTS - PUBLICATIONS**

# DTTC Publications January 2018 - December 2018

- Allwood BW, Maarman GJ, Kyriakakis CG, Doubell, AF. Post-pulmonary tuberculosis complications in South Africa and a potential link with pulmonary hypertension: Premise for clinical and scientific investigations. S Afr Med J. 2018 Jun. 26;108(7):12339. doi: 10.7196/SAMJ.2018.v108i7.13359. PubMed PMID: 30004336.
- 2. Basu-Roy R, Whittaker E, **Seddon JA**, Kampmann B. Tuberculosis susceptibility and protection in children. Lancet Infect Dis. 2018 Oct 12. pii: S1473-3099(18)30157-9. doi: 10.1016/S1473-3099(18)30157-9. Epub ahead of print. PubMed PMID: 30322790.
- 3. **Bock P**, Fatti G, Ford N, Jennings K, Kruger J, Gunst C, Louis F, Grobbelaar N, Shanaube K, Floyd S, Grimwood A, Hayes R, Ayles H, Fidler S, **Beyers N**; HPTN 071 (PopART) trial team. Attrition when providing antiretroviral treatment at CD4 counts >500cells/muL at three government clinics included in the HPTN 071 (PopART) trial in South Africa. PLoS One. 2018 Apr 19;13(4):e0195127. doi: 10.1371/journal.pone.0195127. PubMed PMID: 29663952.
- 4. Bock P, Jennings K, Vermaak R, Cox H, Meintjes G, Fatti G, Kruger J, De Azevedo V, Maschilla L, Louis F, Gunst C, Grobbelaar N, Dunbar R, Limbada M, Floyd S, Grimwood A, Ayles H, Hayes R, Fidler S, Beyers N. Incidence of Tuberculosis among HIV-positive individuals initiating antiretroviral treatment at higher CD4 counts in the HPTN 071 (PopART) trial in South Africa. J Acquir Immune Defic Syndr. 2018 Jan 1;77(1):93-101. doi: 10.1097/QAI.00000000000001560. PubMed PMID: 29016524.
- 5. Bond, V, Ngwenya F, **Thomas A**, Simuyaba M, **Hoddinott G**, Fidler S, Hayes R, Ayles H, Seeley, J; HPTN 071 (PopART) study team. Spinning plates: livelihood mobility, household responsibility and anti-retroviral treatment in an urban Zambian community during the HPTN 071 (PopART) study. J Int AIDS Soc. 2018 Jul;21 Suppl 4:e25117. doi.org/10.1002/jia2.25117. PubMed PMID: 30027643.
- 6. Bond V, Nomsenge S, Mwamba M, Ziba D, Birch A, **Mubekapi-Musadaidzwa C**, **Vanqa N, Viljoen L,** Pliakas T, Ayles H, Hargreaves J, **Hoddinott G**, Stangl A, Seeley J; HPTN 071 (PopART) study team. "Being seen" at the clinic: Zambian and South African health worker reflections on the relationship between health facility spatial organisation and items and HIV stigma in 21 health facilities, the HPTN 071 (PopART) study. Health Place. 2018 Dec 7. pii: S1353-8292(18)30608-7. doi: 10.1016/j.healthplace.2018.11.006. [Epub ahead of print]. PubMed PMID: 30528346.
- 7. Bond V, Ngwenya F, Murray E, Ngwenya N, **Viljoen L,** Gumede D, Bwalya C, **Mantantana J, Hoddinott G**, Dodd PJ, Ayles H, Simwinga M, Wallman S, Seeley J. Value and Limitations of Broad Brush Surveys Used in Community-Randomized Trials in Southern Africa. Qual Health Res. 2018 Dec 17:1049732318809940. doi: 10.1177/1049732318809940. [Epub ahead of print]. PubMed PMID: 30556470.
- 8. Bradley J, Floyd S, Piwowar-Manning E, Laeyendecker O, Young A, **Bell-Mandla N**, Bwalya J, **Bock P**, Fidler S, Ayles H, Hayes, RJ; HPTN 071(PopART) Study Team. Sexually transmitted bedfellows: exquisite association between HIV and Herpes Simplex Virus Type 2 in 21 Communities in Southern Africa in the HIV Prevention Trials Network 071 (PopART) study. J Infect Dis. 2018 Jul 2;218(3):443-452. doi: 10.1093/infdis/jiy178. PubMed PMID: 29659909.
- 9. Byamungu LN, **du Preez K**, **Walters E**, Nachega JB, **Schaaf HS**. Timing of HIV diagnosis in children with tuberculosis managed at a referral hospital in Cape Town, South Africa. Int J Tuberc Lung Dis. 2018 May 1;22(5):488-495. doi: 10.5588/ijtld.17.0613. PubMed PMID: 29663952.
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- SHINE trial team. Shorter treatment for minimal tuberculosis (TB) in children (SHINE): a study protocol for a randomised controlled trial. Trials. 2018 Apr 19;19(1):237. doi: 10.1186/s13063-018-2608-5. PubMed PMID: 29673395.
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- 13. Denti P, **Garcia-Prats AJ**, **Draper HR**, Wiesner L, **Winckler J**, Thee S, Dooley KE, Savic RM, McIlleron HM, **Schaaf HS**, **Hesseling AC**. Levofloxacin population pharmacokinetics in South African children treated for multidrug-resistant tuberculosis. Antimicrob Agents Chemother. 2018 Jan 25;62(2). pii: e01521-17. doi: 10.1128/AAC.01521-17. PubMed PMID: 29133560.
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- 16. du Preez K, Schaaf HS, Dunbar R, Walters E, Swartz A, Solomons R, Hesseling AC. Complementary surveillance strategies are needed to better characterise the epidemiology, care pathways and treatment outcomes of tuberculosis in children. BMC Public Health. 2018 Mar 23;18(1):397. doi: 10.1186/s12889-018-5252-9. PubMed PMID: 29566651.
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- of longitudinal qualitative data from the HPTN 071 (PopART) trial. J Int AIDS Soc. 2018 Jul;21 Suppl 4:e25135. doi: 10.1002/jia2.25135. PubMed PMID: 30027687.
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# **Other Publications**

- Schaaf HS, Marais BJ, Carvalho I, Seddon JA. Challenges in childhood tuberculosis. In: ERS Monograph: *Tuberculosis*. Eds. Migliori GB, Bothamley G, Duarte R, Rendon A. European Respiratory Society, Sheffield, UK. 2018:234-262.
- Statement on injectable-free regimens for children under the age of 12 years with rifampicinresistant tuberculosis <a href="http://sentinel-project.org/wp-content/uploads/2018/07/Recommendations-for-Injectible-Free-Regimens-in-Children-with-Rif-Resistant-TB.pdf">http://sentinel-project.org/wpcontent/uploads/2018/07/Recommendations-for-Injectible-Free-Regimens-in-Children-with-Rif-Resistant-TB.pdf</a>
- Research Priorities for Paediatric Tuberculosis. World Health Organization. <a href="https://www.finddx.org/wp-content/uploads/2018/09/Paediatric">https://www.finddx.org/wp-content/uploads/2018/09/Paediatric</a> TB ResearchPriorities FINAL Web.pdf
- 4. Roadmap towards ending TB in children and adolescents. World Health Organization <a href="http://apps.who.int/iris/bitstream/handle/10665/275422/9789241514798-eng.pdf">http://apps.who.int/iris/bitstream/handle/10665/275422/9789241514798-eng.pdf</a>

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# APPENDIX II: STUDENTS - MASTERS, PHD AND POSTDOCTORAL

# The following PhD students graduated in 2018:

Dr. Anthony Garcia-Prats

Supervisor: Anneke Hesseling; Simon Schaaf

Dissertation: Optimizing and operationalizing MDR-TB treatment in children

Dr. Elisabetta Walters

Supervisors: Anneke Hesseling; Robert Gie

Dissertation: Novel approaches to diagnosis of TB in children

Dr. Peter Bock

Supervisors: Nulda Beyers, Sarah Fidler (Imperial College)

Dissertation: The impact of POPART intervention on the mortality and AIDS related morbidity amongst

HIV positive adults in South Africa

Dr. Florian Marx

Supervisors: Nulda Beyers; Ted Cohen

Dissertations: Mathematical modelling to project the impact of interventions targeted to individuals previously treated for tuberculosis on the trajectory of the tuberculosis epidemic in high-burden settings

#### Sue-Ann Meehan

Supervisors: Nulda Beyers; Ronelle Burger

Dissertation: The contribution of a community based HIV counseling and testing initiative in working

towards increasing access to HIV counseling and testing in Cape Town, South Africa.

Rory Dunbar

Supervisors: Nulda Beyers; Ivor Langley; Pren Naidoo

Dissertation: How can virtual implementation modelling inform the scale-up of new molecular

diagnostic tools for tuberculosis?

#### The following PhD student submitted their dissertations in 2018:

**Graeme Hoddinott** 

Supervisor: Mary van der Riet

Dissertation: Toward a conceptual model of 'the act'; an exercise in theory-generation in the problematic space of school-based HIV prevention through behavior change interventio

| Currently Registered Trainees: Master's Degree |                                                   |                                   |                                                        |                                     |               |                                       |                                                                                                                                    |                                                                                                                                                             |
|------------------------------------------------|---------------------------------------------------|-----------------------------------|--------------------------------------------------------|-------------------------------------|---------------|---------------------------------------|------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Name of trainee                                | Current                                           |                                   | Degree                                                 |                                     | Superviso     | or                                    | Researc                                                                                                                            | th Topic                                                                                                                                                    |
| Jabulile<br>Mantantana,<br>BPsych              | Community<br>Engagement,                          |                                   | MPhil Graeme<br>Transdisciplinary<br>Public health and |                                     |               | Continuit                             | by of ΔRT for neonle who experience incorporation                                                                                  |                                                                                                                                                             |
| Christopher<br>Mahwire, MI                     | Melissa Nel                                       | Rese                              | al Science<br>arch<br>entice                           |                                     |               | Graeme<br>Hoddino                     | tt                                                                                                                                 | A scale for rapidly ascribing socio-economic status estimates to neighbourhoods using observational and participatory research techniques.                  |
| Marcel Kany<br>Kitnge<br>Nozizwe Mal           | Nelis<br>Grobbelaar                               | Progr<br>Direct<br>Healt          | amme<br>tor (ANOVA<br>h)                               | MPhil<br>Transo<br>Public<br>develo |               |                                       | tt                                                                                                                                 | Integrating change to universal HIV-testing and ART regardless of CD4 -count into routine health services                                                   |
| Lena Ronge,                                    | Dianne van Aswegen Social Science Research Intern |                                   | MA Anthropology (SU)                                   |                                     | Shaheed Tayob |                                       | Trauma, care and evidence, from the Mosaic centre to the courtroom, gender-based violence and the state in Cape Town, South Africa |                                                                                                                                                             |
| MD<br>Nosivuyile<br>Vanqa                      | Kyla Meyerson                                     | Social Science<br>Research Intern |                                                        | (SU)                                |               | Mark Tomlinson<br>Graeme<br>Hoddinott |                                                                                                                                    | The experiences of paediatric MDR-TB patients' (0-5 years old) hospitalised for treatment in terms of separation and attachment                             |
|                                                | Leletu Busakwe                                    | Social Science<br>Research Intern |                                                        | MA Anthropology<br>(UWC)            |               | Sakhumzi<br>Mfecane                   |                                                                                                                                    | The experiences of paediatric MDR-TB patients' (14-17 years old) hospitalised for treatment in terms of adolescent social and biological development        |
| Dionne Jivar                                   | Vuyokazi Myoli                                    |                                   | l Science<br>arch Intern                               |                                     |               |                                       |                                                                                                                                    | Young women's talk about reproductive health service access                                                                                                 |
|                                                | Khanya Mama                                       |                                   |                                                        | MCom                                |               | Ronelle I<br>Graeme<br>Hoddino        | _                                                                                                                                  | TB patients with confirmed diagnoses who decline/delay taking up treatment – exploring why                                                                  |
|                                                | Rene Raad                                         |                                   | l Science<br>arch Intern                               | MA An                               | thropology    | Thomas                                | Cousins                                                                                                                            | Termination of pregnancy providers' experiences of stigma and care                                                                                          |
|                                                | Bianca Hamman                                     | Medio<br>micro                    | cal<br>biology                                         | MSc (S                              | SU)           | Mae Nev<br>Marieke                    | vton-Foot<br>vd Zalm                                                                                                               | The role of respiratory co-infections and the microbiome in the clinical presentation and response to treatment in South African children with suspected TB |

| Currently registered PhDs      |                              |              |                                                    |                                                                                                                                                                              |  |
|--------------------------------|------------------------------|--------------|----------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--|
| Name of trainee                | Current<br>position          | Degree       | Supervisor                                         | Research Topic                                                                                                                                                               |  |
| Nomtha Mandla,<br>Physio       | Project Manager:<br>PopArt   | PHD (SU)     | Lungiswa Nkonki<br>Peter Bock                      | Recruitment, participation and retention of research participants in the HPTN 071 Population Cohort, South Africa.                                                           |  |
| Graeme<br>Hoddinott,<br>MSocSc | Lead Social<br>Scientist     | PhD , (UKZN) | Mary Van Der<br>Riet                               | Toward a conceptual model of 'the act'; an exercise in theory-<br>generation in the problematic space of school-based HIV prevention<br>through behavior change intervention |  |
| Martina<br>Mchenga, MPhil      | PhD student                  | PhD (SU)     | Ronelle Burger                                     | Vulnerable households and health: evidence from surveys in South Africa and Malawi"                                                                                          |  |
| Lario Viljoen,<br>MA           | Social Science<br>Researcher | PhD (SU)     | Lindsey Reynolds                                   | Young women's sexual decision-making in the context of earlier ART-access                                                                                                    |  |
| Karen du Preez,<br>MD, MSc     | Research<br>Medical Officer  | PhD (SU)     | Anneke<br>Hesseling<br>Simon Schaaf<br>Pren Naidoo | Complementary surveillance strategies and interventions to inform a tuberculosis care cascade for children                                                                   |  |
| Anne-Marie<br>Demers, MD       | Medical<br>Microbiologist    | PhD (SU)     | Anneke<br>Hesseling<br>Andrew Whitelaw<br>(NHLS)   | Use of routine microbiology data in paediatric TB trials                                                                                                                     |  |
| Muhammad<br>Osman, MD,<br>Msc  | Research<br>Medical Officer  | PhD (SU)     | Anneke<br>Hesseling<br>Pren Naidoo<br>Alex Welte   | TB-associated mortality in South Africa: longitudinal trends and the impact of health system interventions                                                                   |  |
| Tamaryn Jane<br>Nicholson      | Social Science<br>Researcher | PhD (UKZN)   | Mike Quayle<br>Mary van der Riet<br>Orla Muldoon   | HIV stigma and mass media communication messaging                                                                                                                            |  |
| Louvina van der<br>Laan, MD    | Research<br>Medical Officer  | PhD (UCT)    | Paolo Denti<br>Anthony Garcia-<br>Prats            | Pharmacometric modelling as a tool to optimise TB Treatment in children                                                                                                      |  |

| Post docs current       |                 |              |                     |                                                                       |  |
|-------------------------|-----------------|--------------|---------------------|-----------------------------------------------------------------------|--|
| Name of trainee         |                 | Degree       | Supervisor          | Research Topic                                                        |  |
| Marieke van der<br>Zalm | Paediatrician   | Postdoc (SU) | Anneke<br>Hesseling | Lung function and respiratory pathogens in children with suspected TB |  |
| Rosa Sloot              | Biostatistician | Postdoc (SU) | Pren Naidoo         | Epidemiology- HPTN071 (PopART)                                        |  |

| Graduates | 2018 |
|-----------|------|
|-----------|------|

| Name of trainee                                                   | Current position                                      | Degree                  | Supervisor                                             | Research Topic                                                                                                                                                                                    |  |
|-------------------------------------------------------------------|-------------------------------------------------------|-------------------------|--------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--|
| Heidi van Deventer,<br>MD                                         | Research<br>Medical Officer                           | MSc Clin Epid<br>(SU)   | Anneke<br>Hesseling                                    | Effect of TB on risk of atopy in children                                                                                                                                                         |  |
| Celeste de Vaal,<br>MD, DCH                                       | Research<br>Medical Officer                           | MPhil<br>Bioethics (SU) | Lyn Horn                                               | Ethics of post-mortem studies and notifiable diseases including TB                                                                                                                                |  |
| Catherine Wiseman,<br>MD                                          | Research<br>Medical Officer                           | MPH (UCT)               | Anneke<br>Hesseling                                    | Effect of HIV exposure on risk of TB infection in infants in high-<br>burden communities                                                                                                          |  |
| Anelet James, MSc<br>Microbiology                                 | DTTC<br>Operations<br>Manager                         | MBA (USB)               | Jako Volschenk                                         | Comparing patient costs for MDR TB and HIV co-infected patients under LPA and Xpert diagnostic algorithms                                                                                         |  |
| Anthony Garcia-<br>Prats, MD, Msc                                 | Paediatric PI<br>BCH PK Unit<br>Medical Director      | PhD (SU)                | Anneke<br>Hesseling<br>Simon Schaaf                    | Optimizing and operationalizing MDR-TB treatment in children                                                                                                                                      |  |
| Elisabetta Walters,<br>MD, MMed                                   | Research<br>Pediatrician                              | PhD (SU)                | Anneke<br>Hesseling<br>Robert Gie                      | Novel approaches to diagnosis of TB in children                                                                                                                                                   |  |
| Peter Bock, MBChB<br>(UCT), MRCP (UK),<br>MRCGP (UK) MPH<br>(UCT) | Specialist<br>Family<br>Physician<br>Co-PI: PopArt    | PhD (SU)                | Nulda Beyers,<br>Sarah Fidler<br>(Imperial<br>College) | The impact of POPART intervention on the mortality and AIDS related morbidity amongst HIV positive adults in South Africa                                                                         |  |
| Florian Marx, MD*                                                 | Research Fellow                                       | PhD (SU)                | Nulda Beyers<br>Ted Cohen                              | Mathematical modelling to project the impact of interventions targeted to individuals previously treated for tuberculosis on the trajectory of the tuberculosis epidemic in high-burden settings" |  |
| Sue-Ann Meehan,<br>MA (Research<br>Psychology)                    | Senior<br>Researcher,<br>LINKED-in<br>Project Manager | PhD (SU)                | Nulda Beyers<br>Ronelle Burger                         | The contribution of a community based HIV counseling and testing initiative in working towards increasing access to HIV counseling and testing in Cape Town, South Africa.                        |  |

| Registering for PhDs to start in 2019 |                                    |           |                                                        |                                                                                                      |  |
|---------------------------------------|------------------------------------|-----------|--------------------------------------------------------|------------------------------------------------------------------------------------------------------|--|
| Name of                               | Current                            | Degree    | Supervisor                                             | Research Topic                                                                                       |  |
| trainee                               | position                           |           |                                                        |                                                                                                      |  |
| Hanlie Myburgh,<br>MA                 | Social Science<br>Researcher       | PhD (UVA) | Ria Reis<br>Lindsey Reynolds                           | The state and the citizen in the scale-up of HIV services in South Africa                            |  |
| Constance<br>Mubekapi-<br>Musadaidzwa | Social Science<br>Researcher       | PHD (SU)  | Elona Toska<br>Graeme<br>Hoddinott                     | HIV interventions for young people in southern Africa                                                |  |
| Dillon Wademan                        | Social Science<br>Research Officer | PhD (SU)  | Graeme<br>Hoddinott<br>Lindsey Reynolds                | Family chronicity and intergenerational syndemics in the chronic management of TB, HIV, and diabetes |  |
| Laing de Villiers                     | Social Science<br>Research Officer | PhD (SU)  | Leslie Swartz<br>Graeme<br>Hoddinott                   | Identity fluidity on the margins of Cape Winelands communities                                       |  |
| Rozanne<br>Casper                     | Social Science<br>Research Officer | PhD (RU)  | Catriona Macleod<br>Graeme<br>Hoddinott                | Young women's negotiation of gender identity in South Africa                                         |  |
| Angelique<br>Thomas                   | Social Science<br>Research Officer | PhD (UCT) | Lindsey Reynolds<br>Graeme<br>Hoddinott                | Maintaining intimacy and boundaries by women who sell sex in the<br>Western cape                     |  |
| Isabelle de<br>Wandel, MD             | Medical Officer                    | PhD (SU)  | Anneke<br>Hesseling<br>Marieke Van der<br>Zalm         | TB and Lung Health                                                                                   |  |
| Megan Palmer,<br>MD, FCP              | Paediatrician                      | PhD (SU)  | Anneke<br>Hesseling<br>Pierre Goussard<br>James Seddon | Validity of chest radiographic reading methods in children with<br>suspected tuberculosis            |  |
| Jennifer Hughes                       | Research<br>Medical Officer        | PhD (SU)  | Anthony Garcia-<br>Prats<br>Anneke<br>Hesseling        | MDR-TB treatment strategies in adults and children                                                   |  |

# APPENDIX III: MEDIA INTERACTION (TELEVISION, VIDEO, RADIO, PRINTED MEDIA)

The DTTC has been active in media, both in printed and radio. The centre was represented in a video screening at the 22<sup>nd</sup> International AIDS conference in Amsterdam, 2 printed news articles and three radio interviews.

# Video screening:

- "Two Countries, Two Choices": a 30 minute documentary film on TB in which two DTTC research child participants are featured was released by the organisation, AidsFree World, at the 22<sup>nd</sup> International AIDS Conference in Amsterdam. 27 July 2018. Ref: https://vimeo.com/280906730.

#### Newspaper articles:

- "Weerstandige TB: So slag terapie vir kinders", Elsabie Brits, Die Burger. 18 August 2018.
- Latoya Independent Newspaper. Interview related to an article on TB. Anneke Hesseling. 21 August 2018.

#### Radio:

- Elliot Sylvester and Nicola Brand – CapeTalk. Interview with Anneke Hesseling. 22 August 2018.

- Treating MDR-TB in children. Radio Sonder Grense Mari Hudson. Interview with Anneke Hesseling. 29 August 2018.
- Improvements in TB, SA leading the way globally. Cape Talk. Interview with James Seddon, DTTC. 27 September 2018.